

# CURAZZZZI.APP SEQUENCE LISTING

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- <120> Novel Nucleic Acids and Polypeptides and Methods of Use Thereof
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- Val Pro Ser Gln Gln Ala Thr Glu Asp Met Asp Gln Asp Glu Lys Ser 1140 1145 1150
- Phe Trp Leu Ser Gln Ser Asn Ile Pro Ala Leu Ile Lys Tyr His Met 1155 1160 1165
- Leu Leu Gly Thr Tyr Arg Val Ala Asp Leu Gln Thr Leu Ser Ser 1170 1180
- Asp Met Leu Ala Thr Ser Leu Gln Gly Asn Phe Leu His Leu Ala Lys 1185 1190 1195 1200
- Val Asp Gly Asn Ile Thr Ile Glu Gly Ala Ser Ile Val Asp Gly Asp 1205 1210 1215
- Asn Ala Ala Thr Asn Gly Val Ile His Ile Ile Asn Lys Val Leu Val 1220 1230
- Pro Gln Arg Arg Leu Thr Gly Ser Leu Pro Asn Leu Leu Met Arg Leu 1235 1240 1245
- Glu Gln Met Pro Asp Tyr Ser Ile Phe Arg Gly Tyr Ile Ile Gln Tyr 1250 1255 1260
- Pro Asn Asn Asn Ala Ile Glu Asn Tyr Ile Arg Glu Lys Lys Val Leu 1285 1290 1295
- Ser Leu Glu Glu Asp Val Leu Arg Tyr His Val Val Leu Glu Glu Lys 1300 1305 1310
- Leu Leu Lys Asn Asp Leu His Asn Gly Met His Arg Glu Thr Met Leu 1315 1320 1325
- Gly Phe Ser Tyr Phe Leu Ser Phe Phe Leu His Asn Asp Gln Leu Tyr 1330 1340
- Val Asn Glu Ala Pro Ile Asn Tyr Thr Asn Val Ala Thr Asp Lys Gly 1345 1350 1360

۷al	Ile	нis		Leu 1365	Gly	Lys	٧a٦				1.AP Gln			Arg 1375	Cys
Asp	Asn		Asp 1380		Thr	Ile	Ile	Arg 1385	GТу	Arg	Cys		Thr 1390	Cys	Ser
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Asp	Cys	Asp		G]y 1525	Тгр	Arg	Gly		ніs 1530	Cys	Asp	Asn		Thr 1535	Thr
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	G]y L730	Pro	GТу	Pro		Thr 1735	val	Phe	Ala		Leu 1740	Ser	Ala	Ala	Phe
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Asn	Leu		Leu 1780	Ile	Ser	Asn		Thr 1785	Ser	Leu	Gln		G]u 1790	Pro	Ile
٧a٦		Ser 1795	val	Ser	Gln		Thr 1800	∨al	Tyr	Ile		Asn 1805	Lys	Аlа	Lys
	Ile L810	Ser	Ser	Asp		Ile 1815	Ser	Thr	Asn		11e 1820	٧al	His	Ile	Ile
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Tyr	Ile		Phe 1860	Ser	Asn	Leu		G]n 1865	Asp	Ser	Gly		Leu 1870	Ser	۷a٦
Ile		Asp 1875	Pro	Ile	His		Pro L880	Val	Thr	Leu		Trp L885	Pro	Thr	Asp
	А]а L890	Leu	нis	Ala		His L895	Ala	Leu	Pro		G]u L900	Gln	Gln	Asp	Phe
Leu 1909		Asn	Gln		Asn 1910	Lys	Asp	Lys		Lys L915	Glu	Tyr	Leu	Lys	Phe 1920
His	۷al	Ile		Asp 1925	Ala	Lys	val	Leu 1	А]а L930	٧a٦	Asp	Leu	Pro	Thr 1935	Ser
Thr	Ala		Lys 1940	Thr	Leu	Gln	Gly 1	Ser 1945	Glu	Leu	ser		Lys L950	Cys	Gly
Аlа		Arg 1955	Asp	Ile	Gly		Leu L960	Phe	Leu	Asn		G]n L965	Thr	Tyr	Arg
	Val L970	Gln	Arg	Glu		Leu L975	Phe	Asp	Leu		Val 1980	Аlа	Tyr	Glу	Ile
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Thr Thr Phe Asp Ala Ser Gly Glu Cys Gly Ser Cys Val Asn Thr Pro  $2005 \hspace{1cm} 2010 \hspace{1cm} 2015$ 

Ser Cys Pro Arg Trp Ser Lys Pro Lys Gly Val Lys Gln Lys Cys Leu 2020 2025 2030

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Ser Leu Val Ile Gln Ile Pro Arg Cys Cys Lys Gly Tyr Phe Gly Arg 2050 2055 2060

Asp Cys Gln Ala Cys Pro Gly Gly Pro Asp Ala Pro Cys Asn Asn Arg 2065 2070 2075 2080

Gly Val Cys Leu Asp Gln Tyr Ser Ala Thr Gly Glu Cys Lys Cys Asn 2085 2090 2095

Thr Gly Phe Asn Gly Thr Ala Cys Glu Met Cys Trp Pro Gly Arg Phe 2100 2105 2110

Gly Pro Asp Cys Leu Pro Cys Gly Cys Ser Asp His Gly Gln Cys Asp 2115 2120 2125

Asp Gly Ile Thr Gly Ser Gly Gln Cys Leu Cys Glu Thr Gly Trp Thr 2130 2140

Gly Pro Ser Cys Asp Thr Gln Ala Val Leu Pro Ala Val Cys Thr Pro 2145 2150 2155 2160

Pro Cys Ser Ala His Ala Thr Cys Lys Glu Asn Asn Thr Cys Glu Cys 2165 2170 2175

Asn Leu Asp Tyr Glu Gly Asp Gly Ile Thr Cys Thr Val Val Asp Phe 2180 2185 2190

Cys Lys Gln Asp Asn Gly Gly Cys Ala Lys Val Ala Arg Cys Ser Gln 2195 2200 2205

Lys Gly Thr Lys Val Ser Cys Ser Cys Gln Lys Gly Tyr Lys Gly Asp 2210 2215 2220

Gly His Ser Cys Thr Glu Ile Asp Pro Cys Ala Asp Gly Leu Asn Gly 2225 2230 2235 2240

Gly Cys His Glu His Ala Thr Cys Lys Met Thr Gly Pro Gly Lys His 2245 2250 2255

Lys Cys Glu Cys Lys Ser His Tyr Val Gly Asp Gly Leu Asn Cys Glu 2260 2270

Pro Glu Gln Leu Pro Ile Asp Arg Cys Leu Gln Asp Asn Gly Gln Cys 2275 2280 2285

His Ala Asp Ala Lys Cys Val Asp Leu His Phe Gln Asp Thr Thr Val 2290 2295 2300

Gly Val Phe His Leu Arg Ser Pro Leu Gly Gln Tyr Lys Leu Thr Phe 2305 2310 2315 2320

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Ala Gly Trp Leu Glu Thr Gly Arg Val Ala Tyr Pro Thr Ala Phe Ala 2355 2360 2365

CURA2221.APP Ser Gln Asn Cys Gly Ser Gly Val Val Gly Ile Val Asp Tyr Gly Pro 2370 2375 2380 Arg Pro Asn Lys Ser Glu Met Trp Asp Val Phe Cys Tyr Arg Met Lys Gly Ser Ala Gly Leu Phe Gln Gln Leu Ser Ser Arg Pro Cys Ile Ser 2405 2410 2415 Arg Thr Pro Asp 2420 <210> 5 <211> 11158 <212> DNA <213> Homo sapiens ttgagtttc tctagcatga gaagaatttg cgcggcttgc tggggtctgg cgctcgttc 120 gggctgggcg acctttcagc agatgtccc gtcgcgcaat ttcagcttcc gcctcttccc 180 cgagaccgcg cccggggcc ccggggtat ccccgcgccg cccgctcctg gcgacgaagc 240 ggcggggagc agagtggagc ggctgggca ggctggcca ggcgttccgc gtcgcgctgc tgcgggagct 300 cagcgagcgc ctggagcttg tcttcctggt ggatgattcg tccagcgtgg gcgaagtcaa 360 cttccgcagc gagctcatgt tcgtccgcaa gctgctgcc gacttccccg tggtgccac 420 ggccacgcg gtggccatcg tgaccttctc gtccaagaac tacgtggtgc cgcggtcga 480 ttacatctcc acccgccgc cgcgccagca caagtgcgc ctgctcctcc aagagatccc 540 tgccatctcc taccgagtg gcgaccta acccaagaac accttccaac aagaccacaa 600 caattggtct agggtctccc ccattggaat atccatcagt gatgagaaat acaacgtttg 60 tgccatctcc taccgaggtg gcggcaccta caccaagggc gccttccagc aagccgcgca 600 aattcttctt catgctagag aaaactcaac aaaagttgta tttctcatca ctgatggata 660 ttccaatggg ggagacccta gaccaattgc agcgtcactg cgagattcag gagtggagat 720 cttcactttt ggcatatggc aagggaacat tcgagagctg aatgacatgg cttccacccc 780 aaaggaggag cactgttacc tgctacacag ttttgaagaa tttgaggctt tagtcgccct 840 ctgtcatatg ttatttgtag atctaccttc tgggagtttt attcaagatg atatggtcca 900 ctgctcatat ctttgtgatg aaggcaagga ctgctgtgac cgaatgggaa gctgcaaatg 960 tgggaaacac acaggccatt ttgagtgcat ctgtgaaaag gggtataacg ggaaaggtct 1020 gcagtatgac tgcacagttt gcccatcggg gacatacaaa cctgaaggct caccaggagg 1080 aatcagcagt tgcattccat gtcctgatga aaatcacacc tctccacctg gaagcacatc 1140 ccctgaagac tgtgtctgca gagagggata cagggcatct ggccagacct gtgaagttgt 1200 ccactgcct gccctgaagc ctcccgaaaa tggttacttt atccaaaaca cttgcaacaa 1260 ccacttcaat gcagcctgtg gggtccgatg tcaccctgga tttgatcttg tgggaagcag 1320 catcatctta tgtctaccca atggtttgtg gtccggttca gagagctact gcagagtaag 1380 aacatgtcct catctccgcc agccgaaaca tggccacatc agctgttcta caagggaaat 1440 gttatataag acaacatgtt tggttgcctg tgatgaaggg tacaggctag aaggcagtga 1500 taagcttact tgtcaaggaa acagccagtg ggatgggcca gaaccccggt gtgtggagcg 1560 ccactgttcc acctttcaga tgcccaaaga tgtcatcata tcccccaca actgtggcaa 1620 gcagccagcc aaatttggga cgatctgcta tgtaagttgc cgccaagggt tcattttatc 1680 aattggagat gttgctatcg tatacacggc aactgaccta tccggcaacc aggccagctg 1980 cattttccat atcaaggtta ttgatgcaga accacctgtc atagactggt gcagatctcc 2040 acctcccgtc caggtctcgg agaaggtaca tgccgcaagc tgggatgagc ctcagttctc 2100 agacaactca ggtgctgaat tggtcattac cagaagtcat acacaaggag accttttccc 2160 tcaaggggag actatagtac agtatacagc cactgacccc tcaggcaata acaggacatg 2220 tgatătccat attgtcataa aaggttctcc ctgtgaaatt ccattcacac ctgtaaatgg 2280 ggattttata tgcactccag ataatactgg agtcaactgt acattaactt gcttggaggg 2340 ctatgatttc acagaagggt ctactgacaa gtattattgt gcttatgaag atggcgtctg 2400 gaaaccaaca tataccactg aatggccaga ctgtgccagt aagcgttttg caaaccacgg 2460 gttcaagtcc tttgagatgt tctacaaagc agctcgttgt gatgacacag atctgatgaa 2520 gaagttttct gaagcatttg agacgaccct gggaaaaatg gtcccatcat tttgtagtga 2580 tgcagaggac attgactgca gactggagga gaacctgacc aaaaaatatt gcctagaata 2640

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- Thr Tyr Leu Ser Thr Ala Ser Tyr Ser Cys Asp Thr Gly Tyr Ser Leu 1925 1930 1935
- Gln Gly Pro Ser Ile Ile Glu Cys Thr Ala Ser Gly Ile Trp Asp Arg 1940 1945 1950
- Ala Pro Pro Ala Cys His Leu Val Phe Cys Gly Glu Pro Pro Ala Ile 1955 1960 1965
- Lys Asp Ala Val Ile Thr Gly Asn Asn Phe Thr Phe Arg Asn Thr Val 1970 1975 1980
- Thr Tyr Thr Cys Lys Glu Gly Tyr Thr Leu Ala Gly Leu Asp Thr Ile 1985 1990 1995 2000
- Glu Cys Leu Ala Asp Gly Lys Trp Ser Arg Ser Asp Gln Gln Cys Leu 2005 2010 2015
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Gln Ala Phe Arg Val Arg Leu Leu Arg Glu Leu Ser Glu Arg Leu Glu 65 70 75 80

Leu Val Phe Leu Val Asp Asp Ser Ser Ser Val Gly Glu Val Asn Phe 85 90 95

Arg Ser Glu Leu Met Phe Val Arg Lys Leu Leu Ser Asp Phe Pro Val  $100 \hspace{1cm} 105 \hspace{1cm} 110$ 

Val Pro Thr Ala Thr Arg Val Ala Ile Val Thr Phe Ser Ser Lys Asn 115 120 125

Tyr Val Val Pro Arg Val Asp Tyr Ile Ser Thr Arg Arg Ala Arg Gln 130 135 140

His Lys Cys Ala Leu Leu Elu Gln Glu Ile Pro Ala Ile Ser Tyr Arg 145 150 155 160

Gly Gly Gly Thr Tyr Thr Lys Gly Ala Phe Gln Gln Ala Ala Gln Ile 165 170 175

Leu Leu His Ala Arg Glu Asn Ser Thr Lys Val Val Phe Leu Ile Thr 180 185 190

Asp Gly Tyr Ser Asn Gly Gly Asp Pro Arg Pro Ile Ala Ala Ser Leu 195 200 205

Arg Asp Ser Gly Val Glu Ile Phe Thr Phe Gly Ile Trp Gln Gly Asn 210 220

Ile Arg Glu Leu Asn Asp Met Ala Ser Thr Pro Lys Glu Glu His Cys 235 230 235 240

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Lys Gln Met Thr Val Asn Lys Ile Met Val His Ala Asp Tyr Asn Glu 65 70 75 80

Leu His Arg Met Gly Ser Asp Ile Thr Leu Leu Gln Leu His Arg His  $90 \hspace{1.5cm} 95$ 

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1090

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Ile Asn Ala Asp Leu Asn Leu Glu Arg Gly His Ala Gln Lys Asn Glu Page 85 Asn Ala Arg Gln Gln Leu Glu Arg Gln Asn Lys Glu Leu Lys Val Lys 1780 1785 1790 1780

Leu Gln Glu Met Glu Gly Thr Val Lys Ser Lys Tyr Lys Ala Ser Ile

Thr Ala Leu Glu Ala Lys Ile Ala Gln Leu Glu Glu Gln Leu Asp Asn 1810 1815 1820

Glu Thr Lys Glu Arg Gln Ala Ala Cys Lys Gln Val Arg Arg Thr Glu 1825 1830 1835

Lys Lys Leu Lys Asp Val Leu Leu Gln Val Asp Asp Glu Arg Asp 1845 1850 1855

Ala Glu Gln Tyr Lys Asp Gln Ala Asp Lys Ala Ser Thr Arg Leu Lys 1860 1865 1870

Gln Leu Lys Arg Gln Leu Glu Glu Ala Glu Glu Ala Gln Arg Ala 1875 1880 1885

Asn Ala Ser Arg Arg Lys Leu Gln Arg Glu Leu Glu Asp Ala Thr Glu

Thr Ala Asp Ala Met Asn Arg Glu Val Ser Ser Leu Lys Asn Lys Leu 1915

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<213> Homo sapiens

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 Ser Ala

 Arg Ile Arg 35
 Cys Pro Lys Gly Ser Lys Ala Tyr Gly Ser His Cys Tyr

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 Gln Lys Arg Pro Ser Gly Asn Leu Val Ser Val Leu Ser Gly Ala Glu 75

 Gly Ser Phe Val Ser Ser Leu Val Lys Ser Ile Gly Asn Ser Tyr Ser 95

 Tyr Val Trp Ile Gly Leu His Asp Pro Thr Gln Gly Thr Glu Pro Asn 110

 Gly Glu Gly Trp Glu Trp Ser Ser Ser Ser Asp Val Met Asn Tyr Phe Ala

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Cys Leu Glu Ile Asn Pro Cys Leu Glu Asn His Gly Gly Cys Asp Lys
Asn Ala Glu Cys Thr Gly Asp Gly Pro Asn Gln Ala Ala Cys Asn Cys
Beu Pro Ala Tyr Thr Gly Asp Gly Lys Val Cys Thr Leu Ile Asn Val
Cys Leu Thr Lys Asn Gly Gly Cys Ser Glu Phe Ala Ile Cys Asn His

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465 470 475 480 Asn Leu Pro Phe Lys Arg Asn Leu Glu Gly Cys Arg Glu Arg Cys Ser 485 490 495 Leu Val Ile Gln Ile Pro Arg Cys Cys Lys Gly Tyr Phe Gly Arg Asp  $500 \hspace{1.5cm} 505 \hspace{1.5cm} 510$ Cys Gln Ala Cys Pro Gly Gly Pro Asp Ala Pro Cys Asn Asn Arg Gly 515 520 525 Val Cys Leu Asp Gln Tyr Ser Ala Thr Gly Glu Cys Lys Cys Asn Thr 530 540 Gly Phe Asn Gly Thr Ala Cys Glu Met Cys Trp Pro Gly Arg Phe Gly 545 550 555 Pro Asp Cys Leu Pro Cys Gly Cys Ser Asp His Gly Gln Cys Asp Asp 565 570 575 Gly Ile Thr Gly Ser Gly Gln Cys Leu Cys Glu Thr Gly Trp Thr Gly 580 585 590 Pro Ser Cys Asp Thr Gln Ala Val Leu Pro Ala Val Cys Thr Pro Pro 595 600 605 Cys Ser Ala His Ala Thr Cys Lys Glu Asn Asn Thr Cys Glu Cys Asn 610 620 Leu Asp Tyr Glu Gly Asp Gly Ile Thr Cys Thr Val Val Asp Phe Cys 635 640 Lys Gln Asp Asn Gly Gly Cys Ala Lys Val Ala Arg Cys Ser Gln Lys 655 Gly Thr Lys Val Ser Cys Ser Cys Gln Lys Gly Tyr Lys Gly Asp Gly 660 665 670 His Ser Cys Thr Glu Ile Asp Pro Cys Ala Asp Gly Leu Ásn Gly Gly 675 680 685 Cys His Glu His Ala Thr Cys Lys Met Thr Gly Pro Gly Lys His Lys
690 695 700 Cys Glu Cys Lys Ser His Tyr Val Gly Asp Gly Leu Asn Cys Glu Pro 705 710 715 720 Glu Gln Leu Pro Ile Asp Arg Cys Leu Gln Asp Asn Gly Gln Cys His 725 730 735 Ala Asp Ala Lys Cys Val Asp Leu His Phe Gln Asp Thr Thr Val Gly 740 750 Val Phe His Leu Arg Ser Pro Leu Gly Gln Tyr Lys Leu Thr Phe Asp 755 760 765 Lys Ala Arg Glu Ala Cys Ala Asn Glu Ala Ala Thr Met Ala Thr Tyr 770 775 780 Asn Gln Leu Ser Tyr Ala Gln Lys Ala Lys Tyr His Leu Cys Ser Ala 785 790 795 800

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1060

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1065

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20 25 30

Gly Ile His Cys Asp Gln Ala Cys Ser Cys Val His Gly Arg Cys Asn 40 45 Gln Gly Pro Leu Gly Asp Gly Ser Cys Asp Cys Asp Val Gly Trp Arg
50 55 60 Gly Val His Cys Asp Asn Ala Thr Thr Glu Asp Asn Cys Asn Gly Thr
65 70 75 80 Cys His Thr Ser Ala Asn Cys Leu Thr Asn Ser Asp Gly Thr Ala Ser 85 90 95 Cys Lys Cys Ala Ala Gly Phe Gln Gly Asn Gly Thr Ile Cys Thr Ala 100 105 110 Ile Asn Ala Cys Glu Ile Ser Asn Gly Gly Cys Ser Ala Lys Ala Asp  $115 \hspace{1.5cm} 120 \hspace{1.5cm} 125$ Cys Lys Arg Thr Thr Pro Gly Arg Arg Val Cys Thr Cys Lys Ala Gly 130 140 Tyr Thr Gly Asp Gly Ile Val Cys Leu Glu Ile Asn Pro Cys Leu Glu 145 150 155 160 Asn His Gly Gly Cys Asp Lys Asn Ala Glu Cys Thr Gln Thr Gly Pro 165 170 175 Asn Gln Ala Cys Asn Cys Leu Pro Ala Tyr Thr Gly Asp Gly Lys 180 185 190 Val Cys Thr Leu Ile Asn Val Cys Leu Thr Lys Asn Gly Gly Cys Ser 195 200 205 Glu Phe Ala Ile Cys Asn His Thr Gly Gln Val Glu Arg Thr Cys Thr 210 215 220 Cys Lys Pro Asn Tyr Ile Gly Asp Gly Phe Thr Cys Arg Gly Ser Ile 225 230 240 Tyr Gln Glu Leu Pro Lys Asn Pro Lys Thr Ser Gln Tyr Phe Phe Gln 245 250 255 Leu Gln Glu His Phe Val Lys Asp Leu Val Gly Pro Gly Pro Phe Thr 260 265 270 Val Phe Ala Pro Leu Ser Ala Ala Phe Asp Glu Glu Ala Arg Val Lys 275 280 285 Asp Trp Asp Lys Tyr Gly Leu Met Pro Gln Val Leu Arg Tyr His Val 290 295 300 Val Ala Cys His Gln Leu Leu Glu Asn Leu Lys Leu Ile Ser Asn 305 310 315 Ala Thr Ser Leu Gln Gly Glu Pro Ile Val Ile Ser Val Ser Gln Ser 325 330 335 Thr Val Tyr Ile Asn Asn Lys Ala Lys Ile Ile Ser Ser Asp Ile Ile 340 345 350Ser Thr Asn Gly Ile Val His Ile Ile Asp Lys Leu Leu Ser Pro Lys 355 360 365 Page 91

Asn Leu Leu Ile Thr Pro Lys Asp Asn Ser Gly Arg Ile Leu Gln Asn 370 375 380 Leu Thr Thr Leu Ala Thr Asn Asn Gly Tyr Ile Lys Phe Ser Asn Leu 385 390 395 400 Ile Gln Asp Ser Gly Leu Leu Ser Val Ile Thr Asp Pro Ile His Thr 405 410 415Pro Val Thr Leu Phe Trp Pro Thr Asp Gln Ala Leu His Ala Leu Pro 420 425 430 Ala Glu Gln Gln Asp Phe Leu Phe Asn Gln Asp Asn Lys Asp Lys Leu 435 440 445 Lys Glu Tyr Leu Lys Phe His Val Ile Arg Asp Ala Lys Val Leu Ala 450 455 460 Val Asp Leu Pro Thr Ser Thr Ala Trp Lys Thr Leu Gln Gly Ser Glu 465 470 475 480 Leu Ser Val Lys Cys Gly Ala Gly Arg Asp Ile Gly Asp Leu Phe Leu 485 490 495Asn Gly Gln Thr Cys Arg Ile Val Gln Arg Glu Leu Leu Phe Asp Leu 500 510 Gly Val Ala Tyr Gly Ile Asp Cys Leu Leu Ile Asp Pro Thr Leu Gly 515 520 525 Gly Arg Cys Asp Thr Phe Thr Thr Phe Asp Ala Ser Gly Glu Cys Gly 530 540 Ser Cys Val Asn Thr Pro Ser Cys Pro Arg Trp Ser Lys Pro Lys Gly 555 550 560 Val Lys Gln Lys Cys Leu Tyr Asn Leu Pro Phe Lys Arg Asn Leu Glu 565 570 575 Gly Cys Arg Glu Arg Cys Ser Leu Val Ile Gln Ile Pro Arg Cys Cys 580 585 590 Lys Gly Tyr Phe Gly Arg Asp Cys Gln Ala Cys Pro Gly Gly Pro Asp 595 600 605 Ala Pro Cys Asn Asn Arg Gly Val Cys Leu Asp Gln Tyr Ser Ala Thr 610 615 620 Gly Glu Cys Lys Cys Asn Thr Gly Phe Asn Gly Thr Ala Cys Glu Met 625 630 635 640 Cys Trp Pro Gly Arg Phe Gly Pro Asp Cys Leu Pro Cys Gly Cys Ser 655 Asp His Gly Gln Cys Asp Asp Gly Ile Thr Gly Ser Gly Gln Cys Leu 660 665 670 Cys Glu Thr Gly Trp Thr Gly Pro Ser Cys Asp Thr Gln Ala Val Leu 675 680 685 Pro Ala Val Cys Thr Pro Pro Cys Ser Ala His Ala Thr Cys Lys Glu 690 695 700 Page 92

Asn Asn Thr Cys Glu Cys Asn Leu Asp Tyr Glu Gly Asp Gly Ile Thr 705 710 715 720 Cys Thr Val Val Asp Phe Cys Lys Gln Asp Asn Gly Gly Cys Ala Lys 725 730 735 Val Ala Arg Cys Ser Gln Lys Gly Thr Lys Val Ser Cys Ser Cys Gln
740 745 750 Lys Gly Tyr Lys Gly Asp Gly His Ser Cys Thr Glu Ile Asp Pro Cys 755 760 765 Ala Asp Gly Leu Asn Gly Gly Cys His Glu His Ala Thr Cys Lys Met 770 780 Thr Gly Pro Gly Lys His Lys Cys Glu Cys Lys Ser His Tyr Val Gly 785 790 795 800 Asp Gly Leu Asn Cys Glu Pro Glu Gln Leu Pro Ile Asp Arg Cys Leu 805 810 815 Gln Asp Asn Gly Gln Cys His Ala Asp Ala Lys Cys Val Asp Leu His 820 825 830 Phe Gln Asp Thr Thr Val Gly Val Phe His Leu Arg Ser Pro Leu Gly 835 840 845 Gln Tyr Lys Leu Thr Phe Asp Lys Ala Arg Glu Ala Cys Ala Asn Glu 850 855 Ala Ala Thr Met Ala Thr Tyr Asn Gln Leu Ser Tyr Ala Gln Lys Ala 865 870 880 865 Lys Tyr His Leu Cys Ser Ala Gly Trp Leu Glu Thr Gly Arg Val Ala 885 890 895 Tyr Pro Thr Ala Phe Ala Ser Gln Asn Cys Gly Ser Gly Val Val Gly 900 905 910 Ile Val Asp Tyr Gly Pro Arg Pro Asn Lys Ser Glu Met Trp Asp Val 915 920 925 Phe Cys Tyr Arg Met Lys Asp Val Asn Cys Thr Cys Lys Val Gly Tyr 930 940 Val Gly Asp Gly Phe Ser Cys Ser Gly Asn Leu Leu Gln Val Leu Met 945 950 955 960 Ser Phe Pro Ser Leu Thr Asn Phe Leu Thr Glu Val Leu Ala Tyr Ser 965 970 975 Asn Ser Ser Ala Arg Gly Arg Ala Phe Leu Glu His Leu Thr Asp Leu 980 985 990 Ser Ile Arg Gly Thr Leu Phe Val Pro Gln Asn Ser Gly Leu Gly Glu 995 1000 1005 Asn Glu Thr Leu Ser Gly Arg Asp Ile Glu His His Leu Ala Asn Val 1010 1015 1020 Ser Met Phe Phe Tyr Asn Asp Leu Val Asn Gly Thr Thr Leu Gln Thr 1025 1030 1035 1040 Page 93

Arg Leu Gly Ser Lys Leu Leu Ile Thr Ala Ser Gln Asp Pro Leu Gln 1045 1055

Pro Thr Glu Thr Arg Phe Val Asp Gly Arg Ala Ile Leu Gln Trp Asp 1060 1065 1070

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Ala Pro Pro Ala Pro Val Thr Leu Thr His Thr Gly Leu Gly Ala Gly 1090 1095 1100

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Glu Ser Glu Glu Asp Ile Asn Val Ala Ala Leu Gly Lys Gln Gln Pro 1140 1145 1150

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<212> PRT

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35 40 45

Ala Lys Ile Ile Ser Ser Asp Ile Ile Ser Thr Asn Gly Ile Val His 50 55 60

Ile Ile Asp Lys Leu Leu Ser Pro Lys Asn Leu Leu Ile Thr Pro Lys 65 70 75 80

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Asn Gly Tyr Ile Lys Phe Ser Asn Leu Ile Gln Asp Ser Gly Leu Leu  $100 \hspace{1cm} 105 \hspace{1cm} 110$ 

Ser Val Ile Thr Asp Pro Ile His Thr Pro Val Thr Leu Phe Trp Pro 115 120 125

Thr Asp Gln Ala Leu His Ala Leu Pro Ala Glu Gln Gln Asp Phe Leu Page 94 Phe Asn Gln Asp Asn Lys Asp Lys Leu Lys Glu Tyr Leu Lys Phe His 145 150 155 160 Val Ile Arg Asp Ala Lys Val Leu Ala Val Asp Leu Pro Thr Ser Thr 165 170 175 Ala Trp Lys Thr Leu Gln Gly Ser Glu Leu Ser Val Lys Cys Gly Ala 180 185 190 Gly Arg Asp Ile Gly Asp Leu Phe Leu Asn Gly Gln Thr Cys Arg Ile 195 200 205 Val Gln Arg Glu Leu Leu Phe Asp Leu Gly Val Ala Tyr Gly Ile Asp 210 215 220 Cys Leu Leu Ile Asp Pro Thr Leu Gly Gly Arg Cys Asp Thr Phe Thr 225 230 235 240 Thr Phe Asp Ala Ser Gly Glu Cys Gly Ser Cys Val Asn Thr Pro Ser 245 250 255 Cys Pro Arg Trp Ser Lys Pro Lys Gly Val Lys Gln Lys Cys Leu Tyr 260 270 Asn Leu Pro Phe Lys Arg Asn Leu Glu Gly Cys Arg Glu Arg Cys Ser 275 280 285 Leu Val Ile Gln Ile Pro Arg Cys Cys Lys Gly Tyr Phe Gly Arg Asp 290 295 300 Cys Gln Ala Cys Pro Gly Gly Pro Asp Ala Pro Cys Asn Asn Arg Gly 305 310 315 320 Val Cys Leu Asp Gln Tyr Ser Ala Thr Gly Glu Cys Lys Cys Asn Thr 325 330 335 Gly Phe Asn Gly Thr Ala Cys Glu Met Cys Trp Pro Gly Arg Phe Gly 340 345 350 Pro Asp Cys Leu Pro Cys Gly Cys Ser Asp His Gly Gln Cys Asp Asp 355 360 365 Gly Ile Thr Gly Ser Gly Gln Cys Leu Cys Glu Thr Gly Trp Thr Gly 370 375Pro Ser Cys Asp Thr Gln Ala Val Leu Pro Ala Val Cys Thr Pro Pro 385 390 395 400 Cys Ser Ala His Ala Thr Cys Lys Glu Asn Asn Thr Cys Glu Cys Asn 405 410 415 Leu Asp Tyr Glu Gly Asp Gly Ile Thr Cys Thr Val Val Asp Phe Cys 420 425 430 Lys Gln Asp Asn Gly Gly Cys Ala Lys Val Ala Arg Cys Ser Gln Lys 435 440 445 Gly Thr Lys Val Ser Cys Ser Cys Gln Lys Gly Tyr Lys Gly Asp Gly 450 460 His Ser Cys Thr Glu Ile Asp Pro Cys Ala Asp Gly Leu Asn Gly Gly

Cys His Glu His Ala Thr Cys Lys Met Thr Gly Pro Gly Lys His Lys 485 490 495 Cys Glu Cys Lys Ser His Tyr Val Gly Asp Gly Leu Asn Cys Glu Pro 500 510 Glu Gln Leu Pro Ile Asp Arg Cys Leu Gln Asp Asn Gly Gln Cys His 515 520 525 Ala Asp Ala Lys Cys Val Asp Leu His Phe Gln Asp Thr Thr Val Gly 530 540 Val Phe His Leu Arg Ser Pro Leu Gly Gln Tyr Lys Leu Thr Phe Asp 545 550 560 Lys Ala Arg Glu Ala Cys Ala Asn Glu Ala Ala Thr Met Ala Thr Tyr 565 570 575 Asn Gln Leu Ser Tyr Ala Gln Lys Ala Lys Tyr His Leu Cys Ser Ala 580 585 590 Gly Trp Leu Glu Thr Gly Arg Val Ala Tyr Pro Thr Ala Phe Ala Ser 595 600 605 Gln Asn Cys Gly Ser Gly Val Val Gly Ile Val Asp Tyr Gly Pro Arg 610 620 Pro Asn Lys Ser Glu Met Trp Asp Val Phe Cys Tyr Arg Met Lys Asp 625 635 640 Val Asn Cys Thr Cys Lys Val Gly Tyr Val Gly Asp Gly Phe Ser Cys 645 650 655 Ser Gly Asn Leu Leu Gln Val Leu Met Ser Phe Pro Ser Leu Thr Asn 660 665 670 Phe Leu Thr Glu Val Leu Ala Tyr Ser Asn Ser Ser Ala Arg Gly Arg 675 680 685 Ala Phe Leu Glu His Leu Thr Asp Leu Ser Ile Arg Gly Thr Leu Phe 690 695 700 Pro Gln Asn Ser Gly Leu Gly Glu Asn Glu Thr Leu Ser Gly Arg 710 715 720 Asp Ile Glu His His Leu Ala Asn Val Ser Met Phe Phe Tyr Asn Asp 725 730 735 Leu Val Asn Gly Thr Thr Leu Gln Thr Arg Val Gly Ser Lys Leu Leu 740 745 750 Ile Thr Ala Ser Gln Asp Pro Leu Gln Pro Thr Glu Thr Arg Phe Val 755 760 765 Asp Gly Arg Ala Ile Leu Gln Trp Asp Ile Phe Ala Ser Asn Gly Ile 770 780 Ile His Val Ile Ser Arg Pro Leu Lys Ala Pro Pro Ala Pro Val Thr 785 790 795 800 Leu Thr His Thr Gly Leu Gly Ala Gly Ile Phe Phe Ala Ile Ile Leu Page 96

Val Thr Gly Ala Val Ala Leu Ala Ala Tyr Ser Tyr Phe Arg Ile Asn Arg Arg Thr Ble Gly Phe Gln His Phe Glu Ser Glu Glu Asp Ile Asn Val Ala Ala Leu Gly Lys Gln Gln Pro Glu Asn Ile Ser Asn Pro Leu Ser Glu Ser Thr Thr Ser Ala Pro Pro Glu Pro Ser Tyr Asp Pro Phe Resort Thr Asp Ser Glu Glu Glu Arg Gln Leu Glu Gly Asn Asp Pro Leu Arg Thr Leu

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<212> PRT

<213> Homo sapiens

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Leu Ala Ser Thr Glu Ala Phe Ser Arg Phe Glu Thr Ile Leu Glu Asn 515 520 525

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CURA2221.APP Cys Ser His Pro Asp Arg Gly Gly Cys Ser Glu Asn Ala Glu Cys Val 865 870 880 Pro Gly Ser Leu Gly Thr His His Cys Thr Cys His Lys Gly Trp Ser 885 890 895 Gly Asp Gly Arg Val Cys Val Ala Ile Asp Glu Cys Glu Leu Asp Val 900 905 910 Gly Gly Cys His Thr Asp Ala Leu Cys Ser Tyr Val Gly Pro Gly 915 920 925 Gln Ser Arg Cys Thr Cys Lys Leu Gly Phe Ala Gly Asp Gly Tyr Gln 930 935 940 Cys Ser Pro Ile Asp Pro Cys Arg Ala Gly Asn Gly Gly Cys His Gly 945 950 955 960 Leu Ala Thr Cys Arg Ala Val Gly Gly Gly Gln Arg Val Cys Thr Cys 965 970 975 Pro Pro Gly Phe Gly Gly Asp Gly Phe Ser Cys Tyr Gly Asp Ile Phe 980 985 990 Arg Glu Leu Glu Ala Asn Ala His Phe Ser Ile Phe Tyr Gln Trp Leu 995 1000 1005 Lys Ser Ala Gly Ile Thr Leu Pro Ala Asp Arg Arg Val Thr Ala Leu 1010 1015 1020 Val Pro Ser Glu Ala Ala Val Arg Gln Leu Ser Pro Glu Asp Arg Ala 1025 1030 1035 1040 Phe Trp Leu Gln Pro Arg Thr Leu Pro Asn Leu Val Arg Ala His Phe 1045 1050 1055 1045 Leu Gln Gly Ala Leu Phe Glu Glu Glu Leu Ala Arg Leu Gly Gly Gln 1060 1065 1070 Glu Val Ala Thr Leu Asn Pro Thr Thr Arg Trp Glu Ile Arg Asn Ile 1075 1080 1085 Ser Gly Arg Val Trp Val Gln Asn Ala Ser Val Asp Val Ala Asp Leu 1090 1095 1100 Leu Ala Thr Asn Gly Val Leu His Ile Leu Ser Gln Val Leu Leu Pro Pro Arg Gly Asp Val Pro Gly Gly Gln Gly Leu Leu Gln Gln Leu Asp 1125 1130 1135 Leu Val Pro Ala Phe Ser Leu Phe Arg Glu Leu Leu Gln His His Gly 1140 1150 Leu Val Pro Gln Ile Glu Ala Ala Thr Ala Tyr Thr Ile Phe Val Pro Thr Asn Arg Ser Leu Glu Ala Gln Gly Asn Ser Ser His Leu Asp Ala

Asp Thr Val Arg His His Val Val Leu Gly Glu Ala Leu Ser Met Glu

1195

Thr Leu Arg Lys Gly Gly His Arg Asn Ser Leu Leu Gly Pro Ala His 1205 1210 1215

Trp Ile Val Phe Tyr Asn His Ser Gly Gln Pro Glu Val Asn His Val 1220 1225 1230

Pro Leu Glu Gly Pro Met Leu Glu Ala Pro Gly Arg Ser Leu Ile Gly 1235 1240 1245

Leu Ser Gly Val Leu Thr Val Gly Ser Ser Arg Cys Leu His Ser His 1250 1255 1260

Ala Glu Ala Leu Arg Glu Lys Cys Val Asn Cys Thr Arg Arg Phe Arg 1265 1270 1275 1280

Cys Thr Gln Gly Phe Gln Leu Gln Asp Thr Pro Arg Lys Ser Cys Val 1285 1290 1295

Tyr Arg Ser Gly Phe Ser Phe Ser Arg Gly Cys Ser Tyr Thr Cys Ala 1300 1305 1310

Lys Lys Ile Gln Val Pro Asp Cys Cys Pro Gly Phe Phe Gly Thr Leu 1315 1320 1325

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Pro Asn Cys Thr Gly Val Cys Asp Cys Ala His Gly Leu Cys Gln Glu 1380 1385 1390

Gly Leu Gln Gly Asp Gly Ser Cys Val Cys Asn Val Gly Trp Gln Gly 1395 1400 1405

Leu Arg Cys Asp Gln Lys Ile Thr Ser Pro Gln Cys Pro Arg Lys Cys 1410 1420

Asp Pro Asn Ala Asn Cys Val Gln Asp Ser Ala Gly Ala Ser Thr Cys 1425 1430 1435 1440

Ala Cys Ala Ala Gly Tyr Ser Gly Asn Gly Ile Phe Cys Ser Glu Val 1445 1450 1455

Asp Pro Cys Ala His Gly His Gly Gly Cys Ser Pro His Ala Asn Cys 1460 1465 1470

Thr Lys Val Ala Pro Gly Gln Arg Thr Cys Thr Cys Gln Asp Gly Tyr 1475 1480 1485

Met Gly Asp Gly Glu Leu Cys Gln Glu Ile Asn Ser Cys Leu Ile His 1490 1495 1500

His Gly Gly Cys His Ile His Ala Glu Cys Ile Pro Thr Gly Pro Gln 1505 1510 1515 1520

Gln Val Ser Cys Ser Cys Arg Glu Gly Tyr Ser Gly Asp Gly Ile Arg 1525 1530 1535

Thr Cys Glu Leu Leu Asp Pro Cys Ser Lys Asn Asn Gly Gly Cys Ser 1540 1545 1550

Pro Tyr Ala Thr Cys Lys Ser Thr Gly Asp Gly Gln Arg Thr Cys Thr 1555 1560 1565

Cys Asp Thr Ala His Thr Val Gly Asp Gly Leu Thr Cys Arg Ala Arg 1570 1575 1580

Val Gly Leu Glu Leu Leu Arg Asp Lys His Ala Ser Phe Phe Ser Leu 1585 1590 1595 1600

Arg Leu Leu Glu Tyr Lys Glu Leu Lys Gly Asp Gly Pro Phe Thr Ile 1605 1610 1615

Phe Val Pro His Ala Asp Leu Met Ser Asn Leu Ser Gln Asp Glu Leu 1620 1625 1630

Ala Arg Ile Arg Ala His Arg Gln Leu Val Phe Arg Tyr His Val Val 1635 1640 1645

Gly Cys Arg Arg Leu Arg Ser Glu Asp Leu Leu Glu Gln Gly Tyr Ala 1650 1655 1660

Thr Ala Leu Ser Gly His Pro Leu Arg Phe Ser Glu Arg Glu Gly Ser 1665 1670 1675 1680

Ile Tyr Leu Asn Asp Phe Ala Arg Val Val Ser Ser Asp His Glu Ala 1685 1690 1695

Val Asn Gly Ile Leu His Phe Ile Asp Arg Val Leu Leu Pro Pro Glu 1700 1705 1710

Ala Leu His Trp Glu Pro Asp Asp Ala Pro Ile Pro Arg Asn Val 1715 1720 1725

Thr Ala Ala Gln Gly Phe Gly Tyr Lys Ile Phe Ser Gly Leu Leu 1730 1740

Lys Val Ala Gly Leu Leu Pro Leu Leu Arg Glu Ala Ser His Arg Pro 1745 1750 1760

Phe Thr Met Leu Trp Pro Thr Asp Ala Ala Phe Arg Ala Leu Pro Pro 1765 1770 1775

Asp Arg Gln Ala Trp Leu Tyr His Glu Asp His Arg Asp Lys Leu Ala 1780 1785 1790

Ala Ile Leu Arg Gly His Met Ile Arg Asn Val Glu Ala Leu Ala Ser 1795 1800 1805

Asp Leu Pro Asn Leu Gly Pro Leu Arg Thr Met His Gly Thr Pro Ile 1810 1815 1820

Ser Phe Ser Cys Ser Arg Thr Arg Pro Gly Glu Leu Met Val Gly Glu 1825 1830 1835 1840

Asp Asp Ala Arg Ile Val Gln Arg His Leu Pro Phe Glu Gly Gly Leu 1845 1850 1855

Ala Tyr Gly Ile Asp Gln Leu Leu Glu Pro Pro Gly Leu Gly Ala Arg 1860 1865 1870

Cys Asp His Phe Glu Thr Arg Pro Leu Arg Leu Asn Thr Cys Ser Ile 1875 1880 1885

Cys Gly Leu Glu Pro Pro Cys Pro Glu Gly Ser Gln Glu Gln Gly Ser 1890 1895 1900

Pro Glu Ala Cys Trp Arg Phe Tyr Pro Lys Phe Trp Thr Ser Pro Pro 1905 1910 1915 1920

Leu His Ser Leu Gly Leu Arg Ser Val Trp Val His Pro Ser Leu Trp 1925 1930 1935

Gly Arg Pro Gln Gly Leu Gly Arg Gly Cys His Arg Asn Cys Val Thr 1940 1945 1950

Thr Trp Lys Pro Ser Cys Cys Pro Gly His Tyr Gly Ser Glu Cys 1955 1960 1965

Gln Ala Cys Pro Gly Gly Pro Ser Ser Pro Cys Ser Asp Arg Gly Val 1970 1975 1980

Cys Met Asp Gly Met Ser Gly Ser Gly Gln Cys Leu Cys Arg Ser Gly 1985 1990 1995 2000

Phe Ala Gly Thr Ala Cys Glu Leu Cys Ala Pro Gly Ala Phe Gly Pro 2005 2010 2015

His Cys Gln Ala Cys Arg Cys Thr Val His Gly Arg Cys Asp Glu Gly 2020 2025 2030

Leu Gly Gly Ser Gly Ser Cys Phe Cys Asp Glu Gly Trp Thr Gly Pro 2035 2040 2045

Arg Cys Glu Val Gln Leu Glu Leu Gln Pro Val Cys Thr Pro Pro Cys 2050 2060

Ala Pro Glu Ala Val Cys Arg Ala Gly Asn Ser Cys Glu Cys Ser Leu 2065 2070 2075 2080

Gly Tyr Glu Gly Asp Gly Arg Val Cys Thr Val Ala Asp Leu Cys Gln 2085 2090 2095

Asp Gly His Gly Gly Cys Ser Glu His Ala Asn Cys Ser Gln Val Gly 2100 2105 2110

Thr Met Val Thr Cys Thr Cys Leu Pro Asp Tyr Glu Gly Asp Gly Trp 2115 2120 2125

Ser Cys Arg Ala Arg Asn Pro Cys Thr Asp Gly His Arg Gly Gly Cys 2130 2140

Ser Glu His Ala Asn Cys Leu Ser Thr Gly Leu Asn Thr Arg Arg Cys 2145 2150 2155 2160

Glu Cys His Ala Gly Tyr Val Gly Asp Gly Leu Gln Cys Leu Glu Glu 2165 2170 2175

Ser Glu Pro Pro Val Asp Arg Cys Leu Gly Gln Pro Pro Cys His 2180 2185 2190

Ser Asp Ala Met Cys Thr Asp Gln His Phe Gln Glu Lys Arg Ala Gly 2195 2200

- Val Phe His Leu Gln Ala Thr Ser Gly Pro Tyr Gly Leu Asn Phe Ser 2210 2215 2220
- Glu Ala Glu Ala Ala Cys Glu Ala Gln Gly Ala Val Leu Ala Ser Phe 2225 2230 2235 2240
- Pro Gln Leu Ser Ala Ala Gln Gln Leu Gly Phe His Leu Cys Leu Met 2245 2250 2255
- Gly Trp Leu Ala Asn Gly Ser Thr Ala His Pro Val Val Phe Pro Val 2260 2265 2270
- Ala Asp Cys Gly Asn Gly Arg Val Gly Val Val Ser Leu Gly Ala Arg 2275 2280 2285
- Lys Asn Leu Ser Glu Arg Trp Asp Ala Tyr Cys Phe Arg Val Gln Asp 2290 2295 2300
- Val Ala Cys Arg Cys Arg Asn Gly Phe Val Gly Asp Gly Ile Ser Thr 2305 2310 2315 2320
- Cys Asn Gly Lys Leu Leu Asp Val Leu Ala Ala Thr Ala Asn Phe Ser 2325 2330 2335
- Thr Phe Tyr Gly Met Leu Leu Gly Tyr Ala Asn Ala Thr Gln Arg Gly 2340 2345 2350
- Leu Asp Phe Leu Asp Phe Leu Asp Asp Glu Leu Thr Tyr Lys Thr Leu 2355 2360 2365
- Phe Val Pro Val Asn Glu Gly Phe Val Asp Asn Met Thr Leu Ser Gly 2370 2375 2380
- Pro Asp Leu Glu Leu His Ala Ser Asn Ala Thr Leu Leu Ser Ala Asn 2385 2390 2395 2400
- Ala Ser Gln Gly Lys Leu Leu Pro Ala His Ser Gly Leu Ser Leu Ile 2405 2410 2415
- Ile Ser Asp Ala Gly Pro Asp Asn Ser Ser Trp Ala Pro Val Ala Pro 2420 2425 2430
- Gly Thr Val Val Val Ser Arg Ile Ile Val Trp Asp Ile Met Ala Phe 2435 2440 2445
- Asn Gly Ile Ile His Ala Leu Ala Ser Pro Leu Leu Ala Pro Pro Gln 2450 2460
- Pro Gln Ala Val Leu Ala Pro Glu Ala Pro Pro Val Ala Ala Gly Val 2465 2470 2475 2480
- Gly Ala Val Leu Ala Ala Gly Ala Leu Leu Gly Leu Val Ala Gly Ala 2485 2490 2495
- Leu Tyr Leu Arg Ala Arg Gly Lys Pro Thr Gly Phe Gly Phe Ser Ala 2500 2505 2510
- Phe Gln Ala Glu Asp Asp Ala Asp Asp Asp Phe Ser Pro Trp Gln Glu 2515 2520 2525
- Gly Thr Asn Pro Thr Leu Val Ser Val Pro Asn Pro Val Phe Gly Ser 2530 2540

# Asp Thr Phe Cys Glu Pro Phe Asp Asp Ser Leu Leu Glu Glu Asp Phe 2545 2550 2555 2560 Pro Asp Thr Gln Arg Ile Leu Thr Val Lys 2565 2570

<210> 43 <211> 2212 <212> PRT <213> Homo sapiens

<400> 43 Cys Asp Arg Ser Ala Thr Cys Gln Val Thr Ala Asp Gly Lys Thr Ser 1 10 15 Cys Val Cys Arg Glu Ser Glu Val Gly Asp Gly Arg Ala Cys Tyr Gly
20 25 30 His Leu Leu His Glu Val Gln Lys Ala Thr Gln Thr Gly Arg Val Phe 35 40 Leu Gln Leu Arg Val Ala Val Ala Met Met Asp Gln Gly Cys Arg Glu 50 55 60 Ile Leu Thr Thr Ala Gly Pro Phe Thr Val Leu Val Pro Ser Val Ser 65 70 75 80 Ser Phe Ser Ser Arg Thr Met Asn Ala Ser Leu Ala Gln Gln Leu Cys 85 90 95 Arg Gln His Ile Ile Ala Gly Gln His Ile Leu Glu Asp Thr Arg Thr 100 105 110 Gln Gln Thr Arg Arg Trp Trp Thr Leu Ala Gly Gln Glu Ile Thr Val 115 120 125 Phe Asn Gln Phe Thr Lys Tyr Ser Tyr Lys Tyr Lys Asp Gln Pro 130 140 Gin Gln Thr Phe Asn Ile Tyr Lys Ala Asn Asn Ile Ala Ala Asn Gly 145 150 155 160 145 Val Phe His Val Val Thr Gly Leu Arg Trp Gln Ala Pro Ser Gly Thr 165 170 175 Pro Gly Asp Pro Lys Arg Thr Ile Gly Gln Ile Leu Ala Ser Thr Glu 180 185 190 Ala Phe Ser Arg Phe Glu Thr Ile Leu Glu Asn Cys Gly Leu Pro Ser 195 200 205 Ile Leu Asp Gly Pro Gly Pro Phe Thr Val Phe Ala Pro Ser Asn Glu 210 215 220 Ala Val Asp Ser Leu Arg Asp Gly Arg Leu Ile Tyr Leu Phe Thr Ala 225 230 240 Gly Leu Ser Lys Leu Gln Glu Leu Val Arg Tyr His Ile Tyr Asn His 245 250 255 Gly Gln Leu Thr Val Glu Lys Leu Ile Ser Lys Gly Arg Ile Leu Thr 260 265 270 Page 105

Met Ala Asn Gln Val Leu Ala Val Asn Ile Ser Glu Glu Gly Arg Ile 275 280 285 Leu Leu Gly Pro Glu Gly Val Pro Leu Gln Arg Val Asp Val Met Ala 290 295 300 Ala Asn Gly Val Ile His Met Leu Asp Gly Ile Leu Leu Pro Pro Thr 305 310 315 320Ile Leu Pro Ile Leu Pro Lys His Cys Ser Glu Glu Gln His Lys Ile 325 330 335 Val Ala Gly Ser Cys Val Asp Cys Gln Ala Leu Asn Thr Ser Thr Cys 340 345 350 Pro Pro Asn Ser Val Lys Leu Asp Ile Phe Pro Lys Glu Cys Val Tyr 355 360 365 Ile His Asp Pro Thr Gly Leu Asn Val Leu Lys Lys Gly Cys Ala Ser 370 375 380 Tyr Cys Asn Gln Thr Ile Met Glu Gln Gly Cys Cys Lys Gly Phe Phe 385 390 395 400 Gly Pro Asp Cys Thr Gln Cys Pro Gly Gly Phe Ser Asn Pro Cys Tyr 405 415 Gly Lys Gly Asn Cys Ser Asp Gly Ile Gln Gly Asn Gly Ala Cys Leu 420 425 430 Cys Phe Pro Asp Tyr Lys Gly Ile Ala Cys His Ile Cys Ser Asn Pro 435 440 445 Asn Lys His Gly Glu Gln Cys Gln Glu Asp Cys Gly Cys Val His Gly 450 460 Leu Cys Asp Asn Arg Pro Gly Ser Gly Gly Val Cys Gln Gln Gly Thr 465 470 475 480 Cys Ala Pro Gly Phe Ser Gly Arg Phe Cys Asn Glu Ser Met Gly Asp 485 490 495 Cys Gly Pro Thr Gly Leu Ala Gln His Cys His Leu His Ala Arg Cys 500 505 510 Val Ser Gln Glu Gly Val Ala Arg Cys Arg Cys Leu Asp Gly Phe Glu 515 520 525 Gly Asp Gly Phe Ser Cys Thr Pro Ser Asn Pro Cys Ser His Pro Asp 530 540 Arg Gly Gly Cys Ser Glu Asn Ala Glu Cys Val Pro Gly Ser Leu Gly 545 550 560 Thr His His Cys Thr Cys His Lys Gly Trp Ser Gly Asp Gly Arg Val 565 570 575 Cys Val Ala Ile Asp Glu Cys Glu Leu Asp Val Arg Gly Gly Cys His 580 585 590 Thr Asp Ala Leu Cys Ser Tyr Val Gly Pro Gly Gln Ser Arg Cys Thr 595 600 605 Page 106

Cys Lys Leu Gly Phe Ala Gly Asp Gly Tyr Gln Cys Ser Pro Ile Asp 610 620 Pro Cys Arg Ala Gly Asn Gly Gly Cys His Gly Leu Glu Leu Glu Ala 625 630 635 640 Asn Ala His Phe Ser Ile Phe Tyr Gln Trp Leu Lys Ser Ala Gly Ile 645 650 655 Thr Leu Pro Ala Asp Arg Arg Val Thr Ala Leu Val Pro Ser Glu Ala 660 665 670 Ala Val Arg Gln Leu Ser Pro Glu Asp Arg Ala Phe Trp Leu Gln Pro 675 680 685 Arg Thr Leu Pro Asn Leu Val Arg Ala His Phe Leu Gln Gly Ala Leu 690 695 700 Phe Glu Glu Glu Leu Ala Arg Leu Gly Gly Gln Glu Val Ala Thr Leu 705 710 715 720 Asn Pro Thr Thr Arg Trp Glu Ile Arg Asn Ile Ser Gly Arg Val Trp 725 730 735 Val Gln Asn Ala Ser Val Asp Val Ala Asp Leu Leu Ala Thr Asn Gly
740 745 750 Val Leu His Ile Leu Ser Gln Val Leu Leu Pro Pro Arg Gly Asp Val 755 760 765 Pro Gly Gly Gln Gly Leu Leu Gln Gln Leu Asp Leu Val Pro Ala Phe 770 780 Ser Leu Phe Arg Glu Leu Leu Gln His His Gly Leu Val Pro Gln Ile 785 790 795 800 Glu Ala Ala Thr Ala Tyr Thr Ile Phe Val Pro Thr Asn Arg Ser Leu 805 810 815 Glu Ala Gln Gly Asn Ser Ser His Leu Asp Ala Asp Thr Val Arg His 820 825 830 His Val Val Leu Gly Glu Ala Leu Ser Met Glu Thr Leu Arg Lys Gly 835 840 845 Gly His Arg Asn Ser Leu Leu Gly Pro Ala His Trp Ile Val Phe Tyr 850 855 860 Asn His Ser Gly Gln Pro Glu Val Asn His Val Pro Leu Glu Gly Pro 865 870 880 Met Leu Glu Ala Pro Gly Arg Ser Leu Ile Gly Leu Ser Gly Val Leu 885 890 895 Thr Val Gly Ser Ser Arg Cys Leu His Ser His Ala Glu Ala Leu Arg 900 905 910 Glu Lys Cys Val Asn Cys Thr Arg Arg Phe Arg Cys Thr Gln Gly Phe 915 920 925 Gln Leu Gln Asp Thr Pro Arg Lys Ser Cys Val Tyr Arg Ser Gly Phe 930 935 940 Page 107

Ser 945	₽he	Ser	Arg	G1y	Cys 950	Ser	Tyr	Thr	Cys	Ala 955	Lys	Lys	Ile	Gln	va1 960
Pro	Asp	Cys	Cys	Pro 965	Gly	Phe	Phe	Gly	Thr 970	Leu	Cys	Glu	Pro	Cys 975	Pro
GТу	Gly	Leu	G]y 980	Gly	٧a٦	Cys	Ser	Gly 985	His	Gly	Gln	Cys	Gln 990	Asp	Arg
Phe	Leu	G]y 995	Ser	GТу	Glu		His 1000	Cys	His	Glu		Phe .005	His	Gly	Thr
	Cys L010	Glu	Val	Cys		Leu .015	GТу	Arg	Tyr		Pro .020	Asn	Cys	Thr	Gly
Val 1025		Asp	Cys		ніs 1030	Gly	Leu	Cys		G]u .035	Gly	Leu	Gln	Gly 1	Asp .040
Gly	Ser	Cys		Cys .045	Asn	∨al	Gly		G]n .050	Gly	Leu	Arg		Asp .055	Gln
Lys	Ile		Ser L060	Pro	Gln	Cys		Arg .065	Lys	Cys	Asp		Asn .070	Ala	Asn
Cys		G]n L075	Asp	Ser	Ala		Ala .080	Ser	Thr	Cys		Cys .085	Ala	Ala	Gly
	ser LO90	GТу	Asn	Gly		Phe .095	Cys	Ser	Glu		Asp .100	Pro	Cys	Ala	His
Gly 1105		GТу	Gly		Ser L110	Pro	His	Ala		Cys .115	Thr	Lys	val	Ala 1	Pro .120
Gly	Gln	Arg		Cys .125	Thr	Cys	Gln		Gly .130	Tyr	Met	Gly		Gly .135	Glu
Leu	Cys		G]u L140	Ile	Asn	Ser		Leu .145	Ile	ніѕ	His		Gly .150	Cys	His
Ile		Ala 155	Glu	Cys	Ile		Thr .160	Gly	Pro	Gln		va1 .165	Ser	Cys	ser
	Arg L170	Glu	Gly	Tyr		Gly .175	Asp	Gly	Ile		Thr .180	Cys	Glu	Leu	Leu
Asp 1185		Cys	Ser		Asn 190	Asn	Gly	Gly		Ser .195	Pro	Tyr	Ala	Thr 1	Cys 200
Lys	Ser	Thr		Asp .205	Gly	Gln	Arg		Cys .210	Thr	Cys	Asp		Ala 215	His
Thr	٧a٦		Asp 220	Gly	Leu	Thr		Arg .225	Ala	Arg	٧al		Leu 230	Glu	Leu
Leu		Asp L235	Lys	His	Ala		Phe .240	Phe	Ser	Leu		Leu .245	Leu	Glu	Tyr
	G1u L250	Leu	Lys	Gly	Asp 1	G]y .255	Pro	Phe	Thr		Phe .260	۷al	Pro	His	Ala
Asp 1265		Met	Ser		Leu .270	Ser	Gln	Asp	1	Leu .275 .ge 1		Arg	Ile	Arg 1	Ala 280

- His Arg Gln Leu Val Phe Arg Tyr His Val Val Gly Cys Arg Arg Leu 1285 1290 1295
- Arg Ser Glu Asp Leu Leu Glu Gln Gly Tyr Ala Thr Ala Leu Ser Gly 1300 1305 1310
- His Pro Leu Arg Phe Ser Glu Arg Glu Gly Ser Ile Tyr Leu Asn Asp 1315 1320 1325
- Phe Ala Arg Val Val Ser Ser Asp His Glų Ala Val Asn Gly Ile Leu 1330 1340
- His Phe Ile Asp Arg Val Leu Leu Pro Pro Glu Ala Leu His Trp Glu 1345 1350 1355 1360
- Pro Asp Asp Ala Pro Ile Pro Arg Asp Val Thr Ala Ala Ala Gln 1365 1370 1375
- Gly Phe Gly Tyr Lys Ile Phe Ser Gly Leu Leu Lys Val Ala Gly Leu 1380 1385 1390
- Leu Pro Leu Leu Arg Glu Ala Ser His Arg Pro Phe Thr Met Leu Trp 1395 1400 1405
- Pro Thr Asp Ala Ala Phe Arg Ala Leu Pro Pro Asp Arg Gln Ala Trp 1410 1415 1420
- Leu Tyr His Glu Asp His Arg Asp Lys Leu Ala Ala Ile Leu Arg Gly 1425 1430 1435 1440
- His Met Ile Arg Asn Val Glu Ala Leu Ala Ser Asp Leu Pro Asn Leu 1445 1450 1455
- Gly Pro Leu Arg Thr Met His Gly Thr Pro Ile Ser Phe Ser Cys Ser 1460 1465 1470
- Arg Thr Arg Pro Gly Glu Leu Met Val Gly Glu Asp Asp Ala Arg Ile 1475 1480 1485
- Val Gln Arg His Leu Pro Phe Glu Gly Gly Leu Ala Tyr Gly Ile Asp 1490 1495 1500
- Gln Leu Leu Glu Pro Pro Gly Leu Gly Ala Arg Cys Asp His Phe Glu 1505 1510 1515 1520
- Thr Arg Pro Leu Arg Leu Asn Thr Cys Ser Ile Cys Gly Leu Glu Pro 1525 1530 1535
- Pro Cys Pro Glu Gly Ser Gln Glu Gln Gly Ser Pro Glu Ala Cys Trp 1540 1545 1550
- Arg Phe Tyr Pro Lys Phe Trp Thr Ser Pro Pro Leu His Ser Leu Gly 1555 1560 1565
- Leu Arg Ser Val Trp Val His Pro Ser Leu Trp Gly Arg Pro Gln Gly
  1570 1580
- Leu Gly Arg Gly Cys His Arg Asn Cys Val Thr Thr Trp Lys Pro 1585 1590 1595 1600
- Ser Cys Cys Pro Gly His Tyr Gly Ser Glu Cys Gln Ala Cys Pro Gly 1605 1610 1615 Page 109

Gly	Pro		Ser 1620	Pro	Cys	Ser	Asp	Arg 1625	Glу	٧a٦	Cys	Met	Asp 1630	Gly	Met
Ser		Ser 1635	Gly	Gln	Cys		Cys 1640	Arg	Ser	Gly		А]а 1645	G1y	Thr	Αla
	Glu L650	Leu	Cys	Ala		Gly 1655	Аlа	Phe	Gly		ніs 1660	Cys	Gln	Ala	Cys
Arg 1665		Thr	۷a٦		Gly 1670	Arg	Cys	Asp		Gly 1675	Leu	GТу	Gly		Gly 1680
Ser	Cys	Phe		Asp 1685	Glu	Gly	Trp	Thr	Gly 1690	Pro	Arg	Cys		Val 1695	Gln
Leu	Glu		G]n 1700	Pro	val	Cys	Thr	Pro 1705	Pro	Cys	Ala		Glu 1710	Ala	٧a٦
Cys		А7а 1715	Gly	Asn	Ser		G]u 1720	Cys	Ser	Leu		Tyr 1725	Glu	Gly	Asp
	Arg L730	۷a٦	Cys	Thr		А]а 1735	Asp	Leu	Cys		Asp 1740	Gly	His	Gly	Gly
Cys 1745		Glu	His		Asn 1750	Cys	Ser	Gln		Gly 1755	Thr	Met	٧a٦		Cys 1760
Thr	Cys	Leu		Asp 1765	Tyr	Glu	Gly		Gly 1770	Тгр	Ser	Cys		Ala 1775	Arg
Asn	Pro		Thr 1780	Asp	Gly	His	Arg	Gly 1785	Gly	Cys	Ser		ніs 1790	Ala	Asn
Cys		Ser 1795	Thr	Gly	Leu		Thr 1800	Arg	Arg	Cys		Cys L805	His	Аlа	Gly
	val L810	Gly	Asp	Gly		G]n 1815	Cys	Leu	Glu		Ser 1820	Glu	Pro	Pro	۷al
Asp 1825		Cys	Leu		G]n 1830	Pro	Pro	Pro		His 1835	Ser	Asp	Ala		Cys 1840
Thr	Asp	Leu		Phe 1845	G∏n	Glu	Lys		Ala 1850	Gly	٧a٦	Phe		Leu L855	Gln
Ala	Thr		Gly 1860	Pro	Tyr	Gly	Leu 1	Asn L865	Phe	Ser	Glu		G1u L870	Ala	Аlа
Cys		А]а 1875	Gln	Gly	Ala		Leu L880	Ala	Ser	Phe		G]n L885	Leu	Ser	ΑΊа
	G]n L890	Gln	Leu	Gly		His L895	Leu	Cys	Leu		G]y L900	Trp	Leu	Ala	Asn
Gly 1905		Thr	Ala		Pro L910	۷a٦	۷a٦	Phe		Val L915	Ala	Asp	Cys		Asn L920
Gly	Arg	Val		Ile 1925	۷al	Ser	Leu		Ala 1930	Arg	Lys	Asn		Ser L935	Glu

Arg Trp Asp Ala Tyr Cys Phe Arg Val Gln Asp Val Ala Cys Arg Cys 1940 1945 1950

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Arg Asn Gly Phe Val Gly Asp Gly Ile Ser Thr Cys Asn Gly Lys Leu 1955 1960 1965

Leu Asp Val Leu Ala Ala Thr Ala Asn Phe Ser Thr Phe Tyr Gly Met 1970 1975 1980

Leu Leu Gly Tyr Ala Asn Ala Thr Gln Arg Gly Leu Asp Phe Leu Asp 1985 1990 1995 2000

Phe Leu Asp Asp Glu Leu Thr Tyr Lys Thr Leu Phe Val Pro Val Asn 2005 2010 Pro Val Asn

Glu Gly Phe Val Asp Asn Met Thr Leu Ser Gly Pro Asp Leu Glu Leu 2020 2025 2030

His Ala Ser Asn Ala Thr Leu Leu Ser Ala Asn Ala Ser Gln Gly Lys 2035 2040 2045

Leu Leu Pro Ala His Ser Gly Leu Ser Leu Ile Ile Ser Asp Ala Gly 2050 2055 2060

Pro Asp Asn Ser Ser Trp Ala Pro Val Ala Pro Gly Thr Val Val Val 2065 2070 2075 2080

Ser Arg Ile Ile Val Trp Asp Ile Met Ala Phe Asn Gly Ile Ile His 2085 2090 2095

Ala Leu Ala Ser Pro Leu Leu Ala Pro Pro Gln Pro Ala Val Leu Ala 2100 2105 2110

Pro Glu Ala Pro Pro Val Ala Ala Gly Val Gly Ala Val Leu Ala Ala 2115 2120 2125

Gly Ala Leu Leu Gly Leu Val Ala Gly Ala Leu Tyr Leu Arg Ala Arg 2130 2140

Gly Lys Pro Met Gly Phe Gly Phe Ser Ala Phe Gln Ala Glu Asp Asp 2145 2150 2155 2160

Ala Asp Asp Asp Phe Ser Pro Trp Gln Glu Gly Thr Asn Pro Thr Leu 2165 2170 2175

Val Ser Val Pro Asn Pro Val Phe Gly Ser Asp Thr Phe Cys Glu Pro 2180 2185 2190

Phe Asp Asp Ser Leu Leu Glu Glu Asp Phe Pro Asp Thr Gln Arg Ile 2195 2200 2205

Leu Thr Val Lys 2210

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<211> 149

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Fasciclin
Domain Sequence

<400> 44

<210> 45

<211> 149 <212> PRT

<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Fasciclin
 domain sequence

Ala Gly Thr Val Met S Glu Lys Leu Lys Thr Asp Pro Arg Phe Ser Thr Leu Val Ala Ala Leu Glu Ala Ala Asp Leu Val Glu Thr Leu Asn Asn Ser Gly Asp Phe Thr Val Phe Ala Pro Thr Asn Asp Ala Phe Gln Lys Leu Pro Ala Gly Asp Leu Lys Thr Leu Asp Glu Leu Leu Asn Lys Glu Asp Ala Lys Gln Leu Ala Lys Thr Leu Thr Tyr His Val Val Ala Gly 80 Lys Leu Ser Thr Ala Asp Leu Leu Ser Leu Ser Thr Pro Val Leu Thr Ser Leu Gln Gly Ser Lys Ile Thr Val Ser Gly Lys Asn Asp Thr Glu Leu Leu Lys Asp Val Asn Val Leu Lys Val Asn Asn Ala Thr Val Ile

120

Val Glu Ser Asp Ile Glu Thr Thr Asn Gly Val Ile His Val Ile Asp 130 135 140

Arg Val Leu Leu Pro 145

<210> 46

<211> 104

<212> PRT <213> Artificial Sequence

~220<u>></u>

<220>
<223> Description of Artificial Sequence: XLINK domain
 sequence

<400> 46 Gly Glu Val Phe His Tyr Arg Ala Pro Ser Gly Arg Tyr Lys Leu Thr  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Phe Glu Glu Ala Gln Ala Ala Cys Leu Arg Gln Gly Ala Arg Ile Ala 20 25 30

Thr Thr Gly Gln Leu Tyr Ala Ala Trp Lys Gly Gly Phe Asp Gln Cys
35 40 45

Asp Ala Gly Trp Leu Ala Asp Gly Ser Val Arg Tyr Pro Ile Val Lys 50 60

Pro Arg Glu Asn Cys Gly Gly Asp Lys Asp Gly Phe Pro Gly Val Arg 65 70 75 80

Thr Tyr Tyr Leu Phe Pro Asn Gln Thr Gly Phe Pro Asp Asp Pro Ser 85 90 95

Ser Arg Tyr Asp Val Tyr Cys Phe 100

<210> 47

<211> 3567

<212> PRT

<213> Mus musculus

<400> 47

Met Trp Ser Arg Leu Ala Phe Cys Cys Trp Ala Leu Ala Leu Val Ser 1 5 10 15

Gly Trp Thr Asn Phe Gln Pro Val Ala Pro Ser Leu Asn Phe Ser Phe  $20 \hspace{1cm} 25 \hspace{1cm} 30$ 

Arg Leu Phe Pro Glu Ala Ser Pro Gly Ala Leu Gly Arg Leu Ala Val 35 40 45

Pro Pro Ala Ser Ser Glu Glu Glu Ala Ala Gly Ser Lys Val Glu Arg 50 55 60

Leu Gly Arg Ala Phe Arg Ser Arg Val Arg Arg Leu Arg Glu Leu Ser 65 70 75 80

Gly Ser Leu Glu Leu Val Phe Leu Val Asp Glu Ser Ser Ser Val Gly Page 113 Gln Thr Asn Phe Leu Asn Glu Leu Lys Phe Val Arg Lys Leu Leu Ser 100 105 110 Asp Phe Pro Val Val Ser Thr Ala Thr Arg Val Ala Ile Val Thr Phe 115 120 125 Ser Ser Lys Asn Asn Val Val Ala Arg Val Asp Tyr Ile Ser Thr Ser 130 135 140 Arg Ala His Gln His Lys Cys Ala Leu Leu Ser Arg Glu Ile Pro Ala 145 150 150 160 Ile Thr Tyr Arg Gly Gly Gly Thr Tyr Thr Lys Gly Ala Phe Gln Gln 175 Ala Ala Gln Ile Leu Arg His Ser Arg Glu Asn Ser Thr Lys Val Ile 180 185 190 Phe Leu Ile Thr Asp Gly Tyr Ser Asn Gly Gly Asp Pro Arg Pro Ile 195 200 205 Ala Ala Ser Leu Arg Asp Phe Gly Val Glu Ile Phe Thr Phe Gly Ile 210 215 220 Trp Gln Gly Asn Ile Arg Glu Leu Asn Asp Met Ala Ser Thr Pro Lys 235 240 Glu Glu His Cys Tyr Leu Leu His Ser Phe Glu Glu Phe Glu Ala Leu 245 250 255 Ala Arg Arg Ala Leu His Glu Asp Leu Pro Ser Gly Ser Phe Ile Gln 260 265 270 Glu Asp Met Ala Arg Cys Ser Tyr Leu Cys Glu Ala Gly Lys Asp Cys 275 280 285 Cys Asp Arg Met Ala Ser Cys Lys Cys Gly Thr His Thr Gly Gln Phe 290 295 300 Glu Cys Ile Cys Glu Lys Gly Tyr Tyr Gly Lys Gly Leu Gln His Glu 305 310 315 320 Cys Thr Ala Cys Pro Ser Gly Thr Tyr Lys Pro Glu Ala Ser Pro Gly 325 330 335 Gly Ile Ser Thr Cys Ile Pro Cys Pro Asp Val Ser His Thr Ser Pro 340 345 350Pro Gly Ser Thr Ser Pro Glu Asp Cys Val Cys Arg Glu Gly Tyr Gln 355 360 365 Arg Ser Gly Gln Thr Cys Glu Val Val His Cys Pro Ala Leu Lys Pro 370 37.5 380 Pro Glu Asn Gly Phe Phe Ile Gln Asn Thr Cys Lys Asn His Phe Asn 385 390 395 400 Ala Ala Cys Gly Val Arg Cys Arg Pro Gly Phe Asp Leu Val Gly Ser 405 410 415 Ser Ile His Leu Cys Gln Pro Asn Gly Leu Trp Ser Gly Thr Glu Ser

430

Phe Cys Arg Val Arg Thr Cys Pro His Leu Arg Gln Pro Lys His Gly 435 440 445 His Ile Ser Cys Ser Thr Ala Glu Met Ser Tyr Asn Thr Leu Cys Leu 450 455 460 Val Thr Cys Asn Glu Gly Tyr Arg Leu Glu Gly Ser Thr Arg Leu Thr 465 470 475 480 Cys Gln Gly Asn Ala Gln Trp Asp Gly Pro Glu Pro Arg Cys Val Glu 485 490 495 Arg His Cys Ala Thr Phe Gln Lys Pro Lys Gly Val Ile Ile Ser Pro 500 505 510 Pro Ser Cys Gly Lys Gln Pro Ala Arg Pro Gly Met Thr Cys Gln Leu 515 520 525 Ser Cys Arg Gln Gly Tyr Ile Leu Ser Gly Val Arg Glu Val Arg Cys 530 540 Ala Thr Ser Gly Lys Trp Ser Ala Lys Val Gln Thr Ala Val Cys Lys 545 550 555 560 Asp Val Glu Ala Pro Gln Ile Ser Cys Pro Asn Asp Ile Glu Ala Lys 565 570 575 Thr Gly Glu Gln Gln Asp Ser Ala Asn Val Thr Trp Gln Val Pro Thr 580 585 590 Ala Lys Asp Asn Ser Gly Glu Lys Val Ser Val His Val His Pro Ala 595 600 605 Phe Thr Pro Pro Tyr Leu Phe Pro Ile Gly Asp Val Ala Ile Thr Tyr 610 615 620 Thr Ala Thr Asp Ser Ser Gly Asn Gln Ala Ser Cys Thr Phe Tyr Ile 625 630 635 640 Lys Val Ile Asp Val Glu Pro Pro Val Ile Asp Trp Cys Arg Ser Pro 645 650 655 Pro Pro Ile Gln Val Val Glu Lys Glu His Pro Ala Ser Trp Asp Glu
660 665 670 Pro Gln Phe Ser Asp Asn Ser Gly Ala Glu Leu Val Ile Thr Ser Ser 675 680 685 His Thr Gln Gly Asp Met Phe Pro His Gly Glu Thr Val Val Trp Tyr 690 695 700 Thr Ala Thr Asp Pro Ser Gly Asn Asn Arg Thr Cys Asp Ile His Ile 705 710 715 720 Val Ile Lys Gly Ser Pro Cys Glu Val Pro Phe Thr Pro Val Asn Gly
725 730 735 Asp Phe Ile Cys Ala Gln Asp Ser Ala Gly Val Asn Cys Ser Leu Ser 740 750 Cys Lys Glu Gly Tyr Asp Phe Thr Glu Gly Ser Thr Glu Lys Tyr Tyr Page 115

Cys Ala Phe Glu Asp Gly Ile Trp Arg Pro Pro Tyr Ser Thr Glu Trp
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1090

Cys Gly Val Pro Cys Pro Val Gly Glu Phe Ser Arg Ser Gly Leu Thr 1105 1110 1115 1120

1095

Pro Cys Tyr Pro Cys Pro Arg Asp Tyr Tyr Gln Pro Asn Ala Gly Lys 1125 1130 1135

Ser Phe Cys Leu Ala Cys Pro Phe Tyr Gly Thr Thr Thr Ile Thr Gly 1140 1145 1150

Ala Thr Ser Ile Thr Asp Cys Ser Ser Phe Ser Ser Thr Phe Ser Ala 1155 1160 1165

Ala Glu Glu Ser Ile Val Pro Leu Val Ala Pro Gly His Ser Gln Asn 1170 1175 1180

Lys Tyr Glu Val Ser Ser Gln Val Phe His Glu Cys Phe Leu Asn Pro 1185 1190 1195 1200

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Gly Tyr Met Gly Val His Cys Glu Thr Asp Val Asn Glu Cys Gln Ser 1300 1305 1310

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Ser Cys Lys Cys Pro Pro Gly Phe Leu Gly Thr Arg Cys Glu Lys Asn 1330 1335 1340

Val Asp Glu Cys Leu Ser Gln Pro Cys Gln Asn Gly Ala Thr Cys Lys 1345 1350 1355 1360

Asp Gly Ala Asn Ser Phe Arg Cys Gln Cys Pro Ala Gly Phe Thr Gly 1365 1370 1375

Thr His Cys Glu Leu Asn Ile Asn Glu Cys Gln Ser Asn Pro Cys Arg 1380 1385 1390

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Gln Pro Gly Phe Ser Gly His Arg Cys Glu Thr Glu Gln Pro Ser Gly 1410 1415 1420

Phe Asn Leu Asp Phe Glu Val Ser Gly Ile Tyr Gly Tyr Val Leu Leu Page 117 Asp Gly Val Leu Pro Thr Leu His Ala Ile Thr Cys Ala Phe Trp Met

1445 1450 1455

Lys Ser Ser Asp Val Ile Asn Tyr Gly Thr Pro Ile Ser Tyr Ala Leu 1460 1465 1470

Glu Asp Asp Lys Asp Asn Thr Ser Leu Leu Thr Asp Tyr Asn Gly Trp 1475 1480 1485

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Phe Val Gly Ser Ile Ser Gln Leu Asn Leu Trp Asp Tyr Val Leu Ser 1570 1575 1580

Pro Gln Gln Val Lys Leu Leu Ala Ser Ser Cys Pro Glu Glu Leu Ser 1585 1590 1595 1600

Arg Gly Asn Val Leu Ala Trp Pro Asp Phe Leu Ser Gly Ile Thr Gly
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Lys Val Lys Val Asp Ser Ser Ser Met Phe Cys Ser Asp Cys Pro Ser 1620 1625 1630

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Asn Gly Phe Tyr Ser Ala Glu Asp Phe His Ala Gly Ser Thr Val Thr 1700 1705 1710

Tyr Gln Cys Thr Ser Gly Tyr Tyr Leu Leu Gly Asp Ser Arg Met Phe 1715 1720 1725

Cys Thr Asp Asn Gly Ser Trp Asn Gly Ile Ser Pro Ser Cys Leu Asp 1730 1735 1740

Val Asp Glu Cys Ala Val Gly Ser Asp Cys Ser Glu His Ala Ser Cys 1745 1750 1755 1760

Leu Asn Thr Asn Gly Ser Tyr Val Cys Ser Cys Asn Pro Pro Tyr Thr Page 118 Gly Asp Gly Lys Asn Cys Ala Glu Pro Val Lys Cys Lys Ala Pro Glu 1780 1785 1790

Asn Pro Glu Asn Gly His Ser Ser Gly Glu Ile Tyr Thr Val Gly Thr 1795 1800 1805

Ala Val Thr Phe Ser Cys Asp Glu Gly His Glu Leu Val Gly Val Ser 1810 1815 1820

Thr Ile Thr Cys Leu Glu Thr Gly Glu Trp Asp Arg Leu Arg Pro Ser 1825 1830 1835 1840

Cys Glu Ala Ile Ser Cys Gly Val Pro Pro Val Pro Glu Asn Gly Gly 1845 1850 1855

Val Asp Gly Ser Ala Phe Thr Tyr Gly Ser Lys Val Val Tyr Arg Cys 1860 1865 1870

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Ser Gly Ser Trp Ser His Ser Ser Pro Val Cys Glu Leu Val Lys Cys 1890 1895 1900

Ser Gln Pro Glu Asp Ile Asn Asn Gly Lys Tyr Ile Leu Ser Gly Leu 1905 1910 1915 1920

Thr Tyr Leu Ser Ile Ala Ser Tyr Ser Cys Glu Asn Gly Tyr Ser Leu 1925 1930 1935

Gln Gly Pro Ser Leu Leu Glu Cys Thr Ala Ser Gly Ser Trp Asp Arg 1940 1945 1950

Ala Pro Pro Ser Cys Gln Leu Val Ser Cys Gly Glu Pro Pro Ile Val 1955 1960 1965

Lys Asp Ala Val Ile Thr Gly Ser Asn Phe Thr Phe Gly Asn Thr Val 1970 1975 1980

Ala Tyr Thr Cys Lys Glu Gly Tyr Thr Leu Ala Gly Pro Asp Thr Ile 1985 1990 1995 2000

Val Cys Gln Ala Asn Gly Lys Trp Asn Ser Ser Asn His Gln Cys Leu 2005 2010 2015

Ala Val Ser Cys Asp Glu Pro Pro Asn Val Asp His Ala Ser Pro Glu 2020 2025 2030

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Gly Tyr Ser Leu Ala Asp Asn Ser Gln Leu Ile Cys Asn Ala Gln Gly 2050 2060

Asn Trp Val Pro Pro Ala Gly Gln Ala Val Pro Arg Cys Ile Ala His 2065 2070 2075 2080

Phe Cys Glu Lys Pro Pro Ser Val Ser Tyr Ser Ile Leu Glu Ser Val 2085 2090 2095

Ser Lys Ala Lys Phe Ala Ala Gly Ser Val Val Ser Phe Lys Cys Met Page 119

2110

2100

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Gly Glu Trp Ser Pro Ser Pro Leu Ser Val Gln Cys Ile Pro Val Arg 2130 2135 2140

Cys Gly Glu Pro Pro Ser Ile Ala Asn Gly Tyr Pro Ser Gly Thr Asn 2145 2150 2155 2160

Tyr Ser Phe Gly Ala Val Ala Tyr Ser Cys His Lys Gly Phe Tyr 2165 2170 2175

Ile Lys Gly Glu Lys Lys Ser Thr Cys Glu Ala Thr Gly Gln Trp Ser 2180 2185 2190

Lys Pro Thr Pro Thr Cys His Pro Val Ser Cys Asn Glu Pro Pro Lys 2195 2200 2205

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Glu Ala Arg Phe Gln Cys Asn Pro Gly Tyr Lys Ala Ala Gly Ser Pro 2225 2230 2235 2240

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Cys Asn Glu Gly Tyr Glu Leu Val Gly Asp Asn Ser Trp Thr Cys Gln 2290 2295 2300

Lys Ser Gly Lys Trp Ser Lys Lys Pro Ser Pro Lys Cys Val Pro Thr 2305 2310 2315 2320

Lys Cys Ala Glu Pro Pro Leu Leu Glu Asn Gln Leu Val Leu Lys Glu 2325 2330 2335

Leu Ala Ser Glu Val Gly Val Met Thr Ile Ser Cys Lys Glu Gly His 2340 2345 2350

Ala Leu Gln Gly Pro Ser Val Leu Lys Cys Leu Pro Ser Gly Gln Trp 2355 2360 2365

Asn Gly Ser Phe Pro Ile Cys Lys Met Val Leu Cys Pro Ser Pro Pro 2370 2375 2380

Leu Ile Pro Phe Gly Val Pro Ala Ser Ser Gly Ala Leu His Phe Gly 2385 2390 2395 2400

Ser Thr Val Lys Tyr Leu Cys Val Asp Gly Phe Phe Leu Arg Gly Ser 2405 2410 2415

Pro Thr Ile Leu Cys Gln Ala Asp Ser Thr Trp Ser Ser Pro Leu Pro 2420 2425 2430

Glu Cys Val Pro Val Glu Cys Pro Gln Pro Glu Glu Ile Leu Asn Gly Page 120 Ile Ile His Val Gln Gly Leu Ala Tyr Leu Ser Thr Thr Leu Tyr Thr 2450 2455 2460

2440

Cys Lys Pro Gly Phe Glu Leu Val Gly Asn Ala Thr Thr Leu Cys Gly 2465 2470 2475 2480

Glu Asn Gly Gln Trp Leu Gly Gly Lys Pro Met Cys Lys Pro Ile Glu 2485 2490 2495

Cys Pro Glu Pro Lys Glu Ile Leu Asn Gly Gln Phe Ser Ser Val Ser 2500 2505 2510

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Ile Ile Tyr Ser Cys Phe Pro Gly Phe Gln Val Leu Gly His Ala Met 2580 2585 2590

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Lys Val Arg Asp Gly Gln Gly His Phe Asp Gln Glu Asp Asp Met Met 2625 2630 2635 2640

Glu Val Pro Tyr Leu Ala His Pro Gln His Leu Glu Ala Thr Ala Lys 2645 2650 2655

Ala Leu Glu Asn Thr Lys Glu Ser Pro Ala Ser His Ala Ser His Phe 2660 2665 2670

Leu Tyr Gly Thr Met Val Ser Tyr Ser Cys Glu Pro Gly Tyr Glu Leu 2675 2680 2685

Leu Gly Ile Pro Val Leu Ile Cys Gln Glu Asp Gly Thr Trp Asn Gly 2690 2695 2700

Thr Ala Pro Ser Cys Ile Ser Ile Glu Cys Asp Leu Pro Val Ala Pro 2705 2710 2715 2720

Glu Asn Gly Phe Leu His Phe Thr Gln Thr Thr Met Gly Ser Ala Ala 2725 2730 2735

Gln Tyr Ser Cys Lys Pro Gly His Ile Leu Glu Gly Ser His Leu Arg 2740 2745 2750

Leu Cys Leu Gln Asn Lys Gln Trp Ser Gly Thr Val Pro Arg Cys Glu 2755 2760 2765

Ala Ile Ser Cys Ser Lys Pro Asn Pro Leu Trp Asn Gly Ser Ile Lys Page 121 Gly Asp Asp Tyr Ser Tyr Leu Gly Val Leu Tyr Tyr Glu Cys Asp Ser 2785 2790 2795 2800

Gly Tyr Ile Leu Asn Gly Ser Lys Lys Arg Thr Cys Gln Glu Asn Arg 2805 2810 2815

Asp Trp Asp Gly His Glu Pro Met Cys Ile Pro Val Asp Cys Gly Ser 2820 2825 2830

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Lys Leu His Gly Asn Pro Ser Arg Arg Cys Leu Pro Asn Gly Ser Trp 2980 2985 2990

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Ile Gln Gln Gly Thr Ile Asn Ala Thr Asp Leu Gly Cys Gly Lys Thr 3010 3015 3020

Val Gln Ile Glu Cys Phe Lys Gly Phe Lys Leu Leu Gly Leu Ser Glu 3025 3030 3035 3040

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Glu Gly Ser Leu Ser Glu Asp Asn Val Val Thr Tyr Ser Cys Arg Pro 3075 3080 3085

Gly Tyr Thr Met Gln Gly Ser Ser Asp Leu Ile Cys Thr Glu Lys Ala 3090 3095 3100

Ile Trp Ser Gln Pro Tyr Pro Thr Cys Glu Pro Leu Ser Cys Gly Pro Page 122 Pro Pro Thr Val Ala Asn Ala Val Ala Thr Gly Glu Ala His Thr Tyr 3125 3130 3135 3125

Glu Ser Lys Val Lys Leu Arg Cys Leu Glu Gly Tyr Val Met Asp Ser 3145

Asp Thr Asp Thr Phe Thr Cys Gln Gln Asp Gly His Trp Val Pro Glu 3155 3160 3165

Arg Ile Thr Cys Ser Pro Lys Lys Cys Pro Val Pro Ser Asn Met Thr  $3170 \hspace{1cm} 3175 \hspace{1cm} 3180$ 

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Val Ser Cys Ala Glu Gly Phe Thr His Glu Gly Val Asn Trp Ser Thr 3205 3210 3215

Cys Gln Pro Asp Gly Thr Trp Glu Pro Pro Phe Ser Asp Glu Ser Cys

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Val Gly Asn Lys His Ser Phe Gly Ser Thr Ile Val Tyr Gln Cys Asp 3250 3255 3260

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Arg Gln Trp Ser Gly Glu Val Ala Val Cys Arg Glu Asn Arg Cys Glu 3285 3290 3295

Thr Pro Ala Glu Phe Pro Asn Gly Lys Ala Val Leu Glu Asn Thr Thr 3300 3305 3310

Ser Gly Pro Ser Leu Leu Phe Ser Cys His Arg Gly Tyr Thr Leu Glu 3315 3320 3325

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Thr Pro Leu Cys Lys Pro Asn Pro Cys Pro Val Pro Phe Val Ile Pro 3345 3350 3355 3360

Glu Asn Ala Val Leu Ser Glu Lys Glu Phe Tyr Val Asp Gln Asn Val

Ser Ile Lys Cys Arg Glu Gly Phe Leu Leu Lys Gly Asn Gly Val Ile 3380 3385 3390

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Lys Ile Ser Cys Gly Pro Pro Ser His Val Glu Asn Ala Ile Ala Arg 3415 3420

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Gly Tyr Met Leu Glu Gly Ser Leu Arg Ser Val Cys Leu Glu Asn Gly Page 123

Thr Trp Thr Pro Ser Pro Val Cys Arg Ala Val Cys Arg Phe Pro Cys 3465

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Gly Trp Met Gly Arg Leu Cys Glu Glu Pro Ile Cys Ile Leu Pro Cys 3490

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<211> 1316

<212> PRT

<213> Homo sapiens

 CURA2221.APP
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420 425 430 His Ala Ala Ser Trp Asp Glu Pro Gln Phe Ser Asp Asn Ser Gly Ala 435 440 445 Glu Leu Val Ile Thr Arg Ser His Thr Gln Gly Asp Leu Phe Pro Gln 450 460 Gly Glu Thr Ile Val Gln Tyr Thr Ala Thr Asp Pro Ser Gly Asn Asn 465 470 475 480 Arg Thr Cys Asp Ile His Ile Val Ile Lys Gly Ser Pro Cys Glu Ile 485 490 495 Pro Phe Thr Pro Val Asn Gly Asp Phe Ile Cys Thr Pro Asp Asn Thr 500 505 510

Gly Val Asn Cys Thr Leu Thr Cys Leu Glu Gly Tyr Asp Phe Thr Glu 515 520 525 Gly Ser Thr Asp Lys Tyr Tyr Cys Ala Tyr Glu Asp Gly Val Trp Lys 530 540 Pro Thr Tyr Thr Glu Trp Pro Asp Cys Ala Lys Lys Arg Phe Ala 545 550 555 560 Asn His Gly Phe Lys Ser Phe Glu Met Phe Tyr Lys Ala Ala Arg Cys 565 570 575 Asp Asp Thr Asp Leu Met Lys Lys Phe Ser Glu Ala Phe Glu Thr Thr 580 585 590 Leu Gly Lys Met Val Pro Ser Phe Cys Ser Asp Ala Glu Asp Ile Asp 595 600 605 Trp Arg Leu Glu Glu Asn Leu Thr Lys Lys Tyr Cys Leu Glu Tyr Asn 610 620 Tyr Asp Tyr Glu Asn Gly Phe Ala Ile Gly Pro Gly Gly Trp Gly Ala 625 630 635 640 Ala Asn Arg Leu Asp Tyr Ser Tyr Asp Asp Phe Leu Asp Thr Val Gln 645 650 Glu Thr Ala Thr Ser Ile Gly Asn Ala Lys Ser Ser Arg Ile Lys Arg 660 665 670 Ser Ala Pro Leu Ser Asp Tyr Lys Ile Lys Leu Ile Phe Asn Ile Thr 675 680 685 Ala Ser Val Pro Leu Pro Asp Glu Arg Asn Asp Thr Leu Glu Trp Glu 690 700 Asn Gln Gln Arg Leu Leu Gln Thr Leu Glu Thr Ile Thr Asn Lys Leu 705 710 715 720 Lys Arg Thr Leu Asn Lys Asp Pro Met Tyr Ser Phe Gln Leu Ala Ser 725 730 735 Glu Ile Leu Ile Ala Asp Ser Asn Ser Leu Gly Thr Lys Lys Ala Ser 740 745 750 Pro Phe Cys Arg Pro Gly Ser Val Leu Arg Gly Arg Met Cys Val Asn 755 760 765 Cys Pro Leu Gly Thr Tyr Tyr Asn Leu Glu His Phe Thr Cys Glu Ser 770 775 780 Cys Arg Ile Gly Ser Tyr Gln Asp Glu Glu Gly Gln Leu Glu Cys Lys 785 790 795 800 Leu Cys Pro Ser Gly Met Tyr Thr Glu Tyr Ile His Ser Arg Asn Ile 805 810 815 Ser Asp Cys Lys Ala Gln Cys Lys Gln Gly Thr Tyr Ser Cys Ser Gly 820 825 830 Leu Glu Thr Cys Glu Ser Cys Pro Leu Gly Thr Tyr Gln Pro Lys Phe 835 840 845

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Ala Leu Thr Cys Thr Phe Trp Met Lys Ser Ser Asp Asp Met Asn Tyr 1220 1225 1230

Gly Thr Pro Ile Ser Tyr Ala Val Asp Asn Gly Ser Asp Asn Thr Leu 1235 1240 1245

Leu Leu Thr Asp Tyr Asn Gly Trp Val Leu Tyr Val Asn Gly Arg Glu 1250 1255 1260

Lys Ile Thr Asn Cys Pro Ser Val Asn Asp Gly Arg Trp His His Ile 1265 1270 1275 1280

Ala Ile Thr Trp Thr Ser Ala Asn Gly Ile Trp Lys Val Tyr Ile Asp 1285 1290 1295

Gly Lys Leu Ser Asp Gly Gly Ala Gly Leu Ser Val Gly Leu Pro Ile 1300 1305 1310

Pro Gly Met Phe 1315

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<211> 669

<212> PRT

<213> Mus musculus

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Tyr His Gly Gly His Ile Gln Tyr Gln Cys Phe Thr Gly Tyr Lys Leu 65 70 75 80

His Gly Asn Pro Ser Arg Arg Cys Leu Pro Asn Gly Ser Trp Ser Gly  $90 \hspace{1cm} 95$ 

Ser Ser Pro Ser Cys Leu Pro Cys Arg Cys Ser Thr Pro Ile Ile Gln 100 105 110

Gln Gly Thr Ile Asn Ala Thr Asp Leu Gly Cys Gly Lys Thr Val Gln 115 120 125

Ile Glu Cys Phe Lys Gly Phe Lys Leu Leu Gly Leu Ser Glu Ile Thr 130 135 140

Cys Asp Ala Asn Gly Gln Trp Ser Asp Val Pro Leu Cys Glu His Ala 145 150 155 160 Page 128

Gln Cys Gly Pro Leu Pro Thr Ile Pro Asn Ala Ile Val Leu Glu Gly 165 170 175 Ser Leu Ser Glu Asp Asn Val Val Thr Tyr Ser Cys Arg Pro Gly Tyr 180 185 190 Thr Met Gln Gly Ser Ser Asp Leu Ile Cys Thr Glu Lys Ala Ile Trp 195 200 205 Gln Pro Tyr Pro Thr Cys Glu Pro Leu Ser Cys Gly Pro Pro Pro 210 220 Thr Val Ala Asn Ala Val Ala Thr Gly Glu Ala His Thr Tyr Glu Ser 225 230 235 240 Lys Val Lys Leu Arg Cys Leu Glu Gly Tyr Val Met Asp Ser Asp Thr 245 250 255 Asp Thr Phe Thr Cys Gln Gln Asp Gly His Trp Val Pro Glu Arg Ile 260 265 270 Thr Cys Ser Pro Lys Lys Cys Pro Val Pro Ser Asn Met Thr Arg Ile 275 280 285 Arg Phe His Gly Asp Asp Phe Gln Val Asn Arg Gln Val Ser Val Ser 290 295 300 Cys Ala Glu Gly Phe Thr His Glu Gly Val Asn Trp Ser Thr Cys Gln 305 310 315 320 Pro Asp Gly Thr Trp Glu Pro Pro Phe Ser Asp Glu Ser Cys Ile Pro 325 330 335 Val Val Cys Gly His Pro Glu Ser Pro Ala His Gly Ser Val Val Gly 340 345 350 Asn Lys His Ser Phe Gly Ser Thr Ile Val Tyr Gln Cys Asp Pro Gly 355 360 365 Tyr Lys Leu Glu Gly Asn Arg Glu Arg Ile Cys Gln Glu Asn Arg Gln 370 380 Trp Ser Gly Glu Val Ala Val Cys Arg Glu Asn Arg Cys Glu Thr Pro 385 390 395 400 Ala Glu Phe Pro Asn Gly Lys Ala Val Leu Glu Asn Thr Thr Ser Gly 405 410 415 Pro Ser Leu Leu Phe Ser Cys His Arg Gly Tyr Thr Leu Glu Gly Ser 420 425 430 Pro Glu Ala His Cys Thr Ala Asn Gly Thr Trp Asn His Leu Thr Pro 435 440 445 Leu Cys Lys Pro Asn Pro Cys Pro Val Pro Phe Val Ile Pro Glu Asn 450 455 460 Ala Val Leu Ser Glu Lys Glu Phe Tyr Val Asp Gln Asn Val Ser Ile 465 470 475 480 Lys Cys Arg Glu Gly Phe Leu Leu Lys Gly Asn Gly Val Ile Thr Cys 485 490 495 Page 129

 Ser
 Pro
 Asp
 Glu
 Thr
 Trp
 Thr
 His
 Thr
 Asn
 Ala
 Arg
 Cys
 Glu
 Lys
 Ile

 Ser
 Cys
 Gly
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<213> Mus musculus

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Cys Leu Pro Cys Arg Cys Ser Thr Pro Ile Ile Gln Gln Gly Thr Ile
Asn Ala Thr Asp Leu Gly Cys Gly Lys Thr Val Gln Ile Glu Cys Phe
Lys Gly Phe Lys Leu Leu Gly Leu Ser Glu Ile Thr Cys Asp Ala Asn
Gly Gln Trp Ser Asp Val Pro Leu Cys Glu His Ala Gln Cys Gly Pro
Leu Pro Thr Ile Pro Asn Ala Ile Val Leu Glu Gly Ser Leu Ser Glu
Asp Asn Val Val Thr Tyr Ser Cys Arg Pro Gly Tyr Thr Met Gln Gly

Ser Ser Asp Leu Ile Cys Thr Glu Lys Ala Ile Trp Ser Gln Pro Tyr 130 135 140 Pro Thr Cys Glu Pro Leu Ser Cys Gly Pro Pro Pro Thr Val Ala Asn 145 150 155 160 Ala Val Ala Thr Gly Glu Ala His Thr Tyr Glu Ser Lys Val Lys Leu 165 170 175 Lys Cys Leu Glu Gly Tyr Val Met Asp Ser Asp Thr Asp Thr Phe Thr 180 185 190 Cys Gln Gln Asp Gly His Trp Val Pro Glu Arg Ile Thr Cys Ser Pro 195 200 205 Lys Lys Cys Pro Val Pro Ser Asn Met Thr Arg Ile Arg Phe His Gly 210 215 220 Asp Asp Phe Gln Val Asn Arg Gln Val Ser Val Ser Cys Ala Glu Gly 225 230 235 240 Phe Thr His Glu Gly Val Asn Trp Ser Thr Cys Gln Pro Asp Gly Thr 245 250 255 Trp Glu Pro Pro Phe Ser Asp Glu Ser Cys Ile Pro Val Val Cys Gly 260 265 270 His Pro Glu Ser Pro Ala His Gly Ser Val Val Gly Asn Lys His Ser 275 280 285 Phe Gly Ser Thr Ile Val Tyr Gln Cys Asp Pro Gly Tyr Lys Leu Glu 290 295 300 Gly Asn Arg Glu Arg Ile Cys Gln Glu Asn Arg Gln Trp Ser Gly Glu 305 310 315 320 Val Ala Val Cys Arg Glu Asn Arg Cys Glu Thr Pro Ala Glu Phe Pro 325 330 335 Asn Gly Lys Ala Val Leu Glu Asn Thr Thr Ser Gly Pro Ser Leu Leu 340 345 350 Phe Ser Cys His Arg Gly Tyr Thr Leu Glu Gly Ser Pro Glu Ala His 355 360 365 Thr Ala Asn Gly Thr Trp Asn His Leu Thr Pro Leu Cys Lys Pro 370 375 380 Asn Pro Cys Pro Val Pro Phe Val Ile Pro Glu Asn Ala Val Leu Ser 385 390 395 400 Glu Lys Glu Phe Tyr Val Asp Gln Asn Val Ser Ile Lys Cys Arg Glu
405 410 415 Gly Phe Leu Leu Lys Gly Asn Gly Val Ile Thr Cys Ser Pro Asp Glu 420 425 430 Thr Trp Thr His Thr Asn Ala Arg Cys Glu Lys Ile Ser Cys Gly Pro 435 440 445 Pro Ser His Val Glu Asn Ala Ile Ala Arg Gly Val Tyr Tyr Gln Tyr Page 131

450

Val Ala Pro Tyr Gln Cys Asp Cys Pro Thr Gly Trp Thr Gly Ser Arg 545 550 555 560

Cys His Thr Ala Thr Cys Gln Ser Pro Cys Leu Asn Gly Gly Lys Cys 565 570 575

Ile Arg Pro Asn Arg Cys His Cys Leu Ser Ala Trp Thr Gly His Asp 580 585 590

Cys Ser Arg Lys Arg Arg Ala Gly Leu 595 600

<210> 51 <211> 481

<212> PRT <213> Homo sapiens

<400> 51 Met Lys Gly Glu Asn Phe Glu Val Gly Ser Lys Val Gln Phe Phe Cys 1 5 10 15 Asn Glu Gly Tyr Glu Leu Val Gly Asp Ser Ser Trp Thr Cys Gln Lys 20 25 30 Ser Gly Lys Trp Asn Lys Lys Ser Asn Pro Lys Cys Met Pro Ala Lys 40 45Pro Glu Pro Pro Leu Leu Glu Asn Gln Leu Val Leu Lys Glu Leu 50 55 60 Thr Thr Glu Val Gly Val Val Thr Phe Ser Cys Lys Glu Gly His Val 65 70 75 80 Leu Gln Gly Pro Ser Val Leu Lys Cys Leu Pro Ser Gln Gln Trp Asn 85 90 95 Asp Ser Phe Pro Val Cys Lys Ile Val Leu Cys Thr Pro Pro Pro Leu 100 105 110 Ile Ser Phe Gly Val Pro Ile Pro Ser Ser Ala Leu His Phe Gly Ser 115 120 125 Thr Val Lys Tyr Ser Cys Val Gly Gly Phe Phe Leu Arg Gly Asn Ser 130 135 140

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CURA2221.APP Thr Thr Leu Cys Gln Pro Asp Gly Thr Trp Ser Ser Pro Leu Pro Glu 145 150 155 160 Cys Val Pro Val Glu Cys Pro Gln Pro Glu Glu Ile Pro Asn Gly Ile 165 170 175 Ile Asp Val Gln Gly Leu Ala Tyr Leu Ser Thr Ala Leu Tyr Thr Cys 180 185 190 Lys Pro Gly Phe Glu Leu Val Gly Asn Thr Thr Thr Leu Cys Gly Glu 195 200 205 Asn Gly His Trp Leu Gly Gly Lys Pro Thr Cys Lys Ala Ile Glu Cys 210 215 220 Leu Lys Pro Lys Glu Ile Leu Asn Gly Lys Phe Ser Tyr Thr Asp Leu 225 230 235 240 His Tyr Gly Gln Thr Val Thr Tyr Ser Cys Asn Arg Gly Phe Arg Leu 245 250 255 Glu Gly Pro Ser Ala Leu Thr Cys Leu Glu Thr Gly Asp Trp Asp Val 260 265 270 Asp Ala Pro Ser Cys Asn Ala Ile His Cys Asp Ser Pro Gln Pro Ile 275 280 285 Glu Asn Gly Phe Val Glu Gly Ala Asp Tyr Ser Tyr Gly Ala Ile Ile 290 295 300 Ile Tyr Ser Cys Phe Pro Gly Phe Gln Val Ala Gly His Ala Met Gln 305 310 315 320 Thr Cys Glu Glu Ser Gly Trp Ser Ser Ser Ile Pro Thr Cys Met Pro 325 330 335 Ile Asp Cys Gly Leu Pro Pro His Ile Asp Phe Gly Ala Cys Thr Lys 340 345 350Leu Lys Asp Ala Arg Asp Ile Leu Ser Lys Lys Arg His Asp Gly Ser 355 360 365 Ser Ile Cys Asp Ser Ser Pro Ser Leu Ser Phe Gly Ala Val Ala Lys 370 375 380 Thr Trp Glu Asn Thr Lys Glu Ser Pro Ala Thr His Ser Ser Asn Phe 385 390 395 400 Leu Tyr Gly Thr Met Val Ser Tyr Thr Cys Asn Pro Gly Tyr Glu Leu 405 410 415 Leu Gly Asn Pro Val Leu Ile Cys Gln Glu Asp Gly Thr Trp Asn Gly 420 425 430 Ser Ala Pro Ser Cys Ile Ser Ile Glu Cys Asp Leu Pro Thr Ala Pro 435 440 445 Glu Asn Gly Phe Leu Arg Phe Thr Glu Thr Ser Met Gly Ser Ala Val 450 455 460 Gln Tyr Ser Cys Lys Pro Gly His Ile Leu Ala Gly Ser Asp Leu Arg 465 470 475 480 Leu

<220>

domain sequence

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<210> 52
<211> 200
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Von
      willebrand Factor Type A domain sequence
Asp Ile Val Phe Leu Leu Asp Gly Ser Gly Ser Ile Gly Ser Gln Asn 10 15
Phe Glu Arg Val Lys Asp Phe Val Glu Arg Val Val Glu Arg Leu Asp
20 25 30
Val Gly Pro Arg Asp Lys Lys Glu Glu Asp Ala Val Arg Val Gly Leu 35 40 45
Val Gln Tyr Ser Asp Asn Val Arg Thr Glu Ile Lys Phe Lys Leu Asn 50 55 60
Asp Tyr Gln Asn Lys Asp Glu Val Leu Gln Ala Leu Gln Lys Ile Arg 65 70 75 80
Tyr Glu Asp Tyr Tyr Gly Gly Gly Gly Thr Asn Thr Gly Ala Ala Leu
85 90 95
Gln Tyr Val Val Arg Asn Leu Phe Thr Glu Ala Ser Gly Ser Arg Ile
100 105 110
Glu Pro Val Ala Glu Glu Gly Ala Pro Lys Val Leu Val Val Leu Thr
115 120 125
Asp Gly Arg Ser Gln Asp Asp Pro Ser Pro Thr Ile Asp Ile Arg Asp 130 135
Val Leu Asn Glu Leu Lys Lys Glu Ala Gly Val Glu Val Phe Ala Ile
145 150 155 160
Gly Val Gly Asn Ala Asp Asn Asn Asn Leu Glu Glu Leu Arg Glu Ile
165 170 175
Ala Ser Lys Pro Asp Asp His Val Phe Lys Val Ser Asp Phe Glu Ala
                                    185
Leu Asp Thr Leu Gln Glu Leu Leu
<210> 53
<211> 147
<212> PRT
<213> Artificial Sequence
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<223> Description of Artificial Sequence: Pentaxin

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```
Ser Tyr Ala Thr Lys Lys Pro Leu Lys Asp Asn Glu Leu Leu Ile Phe 1 \hspace{1cm} 5 \hspace{1cm} 15
Lys Glu Lys Asp Gly Gln Tyr Ser Leu Tyr Val Gly Gly Ala Pro Gln
20 25 30
Leu Glu Val Thr Phe Lys Val Lys Glu Glu Phe Val Ala Pro Val His
Ile Cys Thr Ser Trp Glu Ser Ser Gly Ile Ala Glu Phe Trp Val 50 60
Asp Gly Lys His Cys Pro Trp Val Arg Lys Gly Leu Lys Lys Gly Tyr 65 70 75 80
Thr Val Gly Ala Glu Pro Ser Ile Ile Leu Gly Gln Glu Gln Asp Ser 85 90 95
Tyr Gly Gly Phe Asp Lys Ser Gln Ser Leu Val Gly Glu Ile Gly 100 105 110
Asp Leu Asn Met Trp Asp Tyr Val Leu Thr Pro Glu Glu Ile Lys Thr
115 120 125
Val Tyr Lys Gly Ala Gly Pro Leu Glu Arg His Ile Tyr Pro Asn Ile
130 135 140
Leu Asp Trp
145
<210> 54
<211> 62
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Sushi domain
       sequence
Cys Pro Pro Pro Asp Ile Glu Asn Gly Arg Val Ser Ser Ser Gly Thr
Tyr Glu Tyr Pro Val Gly Asp Thr Val Thr Tyr Thr Cys Asn Glu Gly
20 25 30
Tyr Arg Leu Val Gly Ser Ser Ser Ile Thr Cys Thr Glu Asp Gly Gly 35 40 45
Gly Gly Trp Ser Pro Pro Leu Leu Gly Glu Leu Pro Lys Cys 50 60
<210> 55
<211> 207
<212> PRT
<213> Homo sapiens
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Gln Leu Val Ala Ala Leu Glu Arg Gln Val Phe Asp Phe Leu Gly Tyr Gln Trp Ala Pro Ile Leu Ala Asn Phe Val His Ile Ile Ile Val Ile Leu Gly Leu Phe Gly Thr Ile Gln Tyr Arg Leu Arg Tyr Val Met Tyr Arg Leu Trp Ala Ala Val Trp Val Thr Trp Asn Val Phe Ile Ile Cys Ro Phe Tyr Leu Glu Val Gly Gly Leu Leu Lys Asp Ser Glu Leu Leu His Gly Thr Phe Ser Leu Ser Arg His Arg Ser Trp Trp Arg Glu Arg Trp Pro Gly Ilo Phe Ilis Glu Glu Val Pro Ala Val Gly Leu Gly Leu Gly Ala Pro His Gly Gly Leu Leu Gly Ala Pro His Gly Gly Leu Gly Ala Leu Gly Pro Ser Tyr Val Ala Leu His Ser Gly Ala Gly Cys Ala Leu Gly Pro Ser Tyr Val Glu Ala Leu His Ser Cys Leu Gln Ile Leu Ile Ala Leu Leu Gly Phe Ilo Cys Gly Cys Gln Val Val Ser Val Phe Thr Glu Glu Glu Asp Ser Phe Asp Phe Ile Gly Gly Phe Asp Phe Asp Phe Ile Gly Gly Phe Asp Pro Leu Tyr His Val Asn Glu Lys Pro Ser Ser Leu Leu Ser Lys Gln Val Tyr Leu Pro Ala

<sup>&</sup>lt;210> 56 <211> 208 <212> PRT

<sup>&</sup>lt;213> Mus musculus

Gly Cys Leu His Glu Glu Glu Ala Thr Ala Gly Leu Gly Ala Leu His 115 120 125 Gly Gln Ser Leu Val Val Gly Ala Gly Cys Ala Met Val His Ser Tyr 130 135 140 Val Glu Ala Leu His Ser Gly Leu Gln Ile Leu Leu Ala Leu Leu Gly 145 150 160 Phe Val Tyr Gly Cys Tyr Val Val Ser Val Leu Thr Glu Glu Asp 165 170 175 Ser Phe Asp Phe Ile Gly Gly Phe Asp Pro Phe Pro Leu Tyr His Val Asn Glu Lys Pro Ser Ser Leu Leu Ser Lys Gln Ala Tyr Leu Pro Ala 195 200 205

<210> 57 <211> 208 <212> PRT

<213> Mus musculus

<400> 57 Met Gly Phe Cys Ser Gly Arg Cys Thr Leu Leu Ala Leu Cys Arg Leu 1 10 15 Gln Leu Val Thr Ala Leu Glu Arg Gln Val Phe Asp Phe Leu Gly Tyr 20 25 30 Gln Trp Ala Pro Ile Leu Ala Asn Phe Thr His Ile Ile Val Val Ile  $35 \hspace{1cm} 40 \hspace{1cm} 45$ Leu Gly Leu Phe Gly Thr Ile Gln Tyr Arg Pro Arg Tyr Ile Val Val 50 55 60 Tyr Val Val Trp Ala Ala Val Trp Val Thr Trp Asn Val Phe Ile Ile 65 70 75 80 Cys Phe Tyr Leu Glu Val Gly Gly Leu Ser Lys Asp Ser Glu Leu Leu 85 90 95 Thr Phe Asn Leu Ser Gly His Arg Ser Trp Trp Glu Glu His Gly Pro 100 105 110Gly Cys Leu His Glu Glu Glu Ala Thr Ala Gly Leu Gly Ala Leu His 115 120 125 Gly Gln Ser Leu Val Val Gly Ala Gly Cys Ala Met Val His Ser Tyr 130 140 Val Glu Ala Leu His Ser Gly Leu Gln Ile Leu Leu Ala Leu Leu Gly 145 150 160 Phe Val Tyr Gly Cys Tyr Val Val Ser Val Leu Thr Glu Glu Asp 165 170 175 Ser Phe Asp Phe Ile Gly Gly Phe Asp Pro Phe Pro Leu Tyr His Val 180

Asn Glu Lys Pro Ser Ser Leu Leu Ser Lys Gln Ala Tyr Leu Pro Ala 200

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<210> 58
<211> 208
<212> PRT
<213> Mus musculus
<400> 58
Met Gly Phe Cys Ser Gly Arg Cys Thr Leu Leu Ala Leu Cys Ala Leu
1 10 15
Gln Leu Val Thr Ala Leu Glu Arg Gln Val Phe Asp Phe Leu Gly Tyr
20 25 30
Gln Trp Ala Pro Ile Leu Ala Asn Phe Thr His Ile Ile Val Val Ile 35 40 45
Leu Gly Leu Phe Gly Thr Ile Gln Tyr Arg Pro Arg Tyr Ile Val Val 50 60
Tyr Val Val Trp Ala Ala Val Trp Val Thr Trp Asn Val Phe Ile Ile 65 70 75 80
Cys Phe Tyr Leu Glu Val Gly Gly Leu Ser Lys Asp Ser Glu Leu Leu 85 90 95
Thr Phe Asn Leu Ser Gly His Arg Ser Trp Trp Glu Glu His Gly Pro
100 105 110
Gly Cys Leu His Glu Glu Glu Ala Thr Ala Gly Leu Gly Ala Leu His
115 120 125
Gly Gln Ser Leu Val Val Gly Ala Gly Cys Ala Met Val His Ser Tyr
130 135 140
Val Glu Ala Leu His Ser Gly Leu Gln Ile Leu Leu Ala Leu Leu Gly
145 150 155 160
Phe Val Tyr Gly Cys Tyr Val Val Arg Val Leu Thr Glu Glu Glu Asp
165 170 175
Ser Phe Asp Phe Ile Gly Gly Phe Asp Pro Phe Pro Leu Tyr His Val
180 185 190
Asn Glu Lys Pro Ser Ser Leu Leu Ser Lys Gln Ala Tyr Leu Pro Ala
195 200 205
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<sup>&</sup>lt;210> 59 <211> 207 <212> PRT <213> Mus musculus

<210> 60 <211> 367

<212> PRT <213> Mus musculus

Met Trp Gly Ser Arg Ala Gln Gln Ser Gly Pro Asp Arg Gly Gly Ala

Cys Leu Leu Ala Ala Phe Leu Leu Cys Phe Ser Leu Leu His Ala Gln

25 Phe Ser Leu Leu His Ala Gln

Asp Tyr Thr Pro Ser Gln Thr Pro Pro Pro Thr Ser Asn Thr Ser Leu

40 45 Pro Arg Gly Arg Val Gln Lys Glu Leu Cys Gly Lys Thr Lys Phe

Gln Gly Lys Ile Tyr Gly Gly Gln Ile Ala Lys Ala Glu Arg Trp Pro 65 70 75 80

CURA2221.APP Trp Gln Ala Ser Leu Ile Phe Arg Gly Arg His Ile Cys Gly Ala Val 85 90 95 Leu Ile Asp Lys Thr Trp Leu Leu Ser Ala Ala His Cys Phe Gln Arg 100 105 110 Ser Leu Thr Pro Ser Asp Tyr Arg Ile Leu Leu Gly Tyr Asn Gln Leu 115 120 125 Ser Asn Pro Ser Asn Tyr Ser Arg Gln Met Thr Val Asn Lys Val Ile 130 135 140 Leu His Glu Asp Tyr Ser Lys Leu Ser Arg Leu Glu Lys Asn Ile Val 145 150 155 160 Leu Ile Gln Leu His His Pro Val Ile Tyr Ser Thr His Ile Phe Pro 165 170 175 Ala Cys Val Pro Asp Gly Thr Thr Lys Val Ser Pro Asn Asn Leu Cys 180 185 190 Trp Ile Ser Gly Trp Gly Met Leu Ser Ala Asp Lys Phe Leu Gln Ala 195 200 205 Pro Phe Pro Leu Leu Asp Ala Glu Val Ser Leu Ile Asp Glu Glu Glu 210 215 220 Cys Thr Thr Phe Phe Gln Thr Pro Glu Val Ser Ile Thr Glu Tyr Asp 225 230 235 240 Val Ile Lys Asp Asp Val Leu Cys Ala Gly Asp Leu Thr Asn Gln Lys 245 250 255 Ser Ser Cys Arg Gly Asp Ser Gly Gly Pro Leu Val Cys Phe Leu Asn 260 270 Ser Phe Trp Tyr Val Val Gly Leu Ala Asn Trp Asn Gly Ala Cys Leu 275 280 285 Glu Pro Ile His Ser Pro Asn Ile Phe Thr Lys Val Ser Tyr Phe Ser 290 295 300 Asp Trp Ile Lys Gln Lys Lys Ala Asn Thr Pro Ala Ala Asp Val Ser 305 310 315 320 Ser Ala Pro Leu Glu Glu Met Ala Ser Ser Leu Arg Gly Trp Gly Asn 325 330 335 Tyr Ser Ala Gly Ile Thr Leu Lys Pro Arg Ile Ser Thr Thr Leu Leu 340 345 350 Ser Ser Gln Ala Leu Leu Leu Gln Ser Ile Trp Leu Arg Ile Leu 355 360 365

<sup>&</sup>lt;210> 61 <211> 366

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Mus musculus

Giy Leu Leu Ala Ala Leu Leu Gly Val Ser Phe Leu Ser Gln His Ala 20 25 30 Gln Thr Ala Glu Pro Thr Asn Val Thr Asn Ala Ala Asn Asn Thr Thr 35 40 45 Ile Gln Ile Met Lys Ser Thr Leu Ser Leu Ser Glu Val Cys Gly Lys 50 60 Thr Lys Phe Gln Gly Lys Ile Tyr Gly Gly Gln Ile Ala Gly Ala Glu 65 70 75 80 Gly Ala Val Leu Ile Asp Lys Asn Trp Val Leu Gly Ala Ala His Cys 100 105 110 Phe Gln Arg Ser Gln Glu Pro Ser Asp Tyr His Val Met Leu Gly Tyr 115 120 125 Thr Asp Leu Asn Ser Pro Thr Arg Tyr Ser Arg Thr Met Ser Val Gln
130 135 140 Lys Val Ile Val His Lys Asp Tyr Asn Arg Phe His Thr Gln Gly Ser 145 150 155 160 Asp Ile Val Leu Leu Gln Leu Arg Ser Ser Val Glu Tyr Ser Ser His 165 170 175 Ile Leu Pro Ala Cys Val Pro Glu Glu Asn Ile Lys Ile Pro Lys Glu 180 185 190 Lys Ala Cys Trp Ala Ser Gly Trp Gly Tyr Leu Arg Glu Asp Val Arg 195 200 205 Ile Pro Leu Pro Asn Glu Leu Tyr Glu Ala Glu Leu Ile Ile Met Ser 210 215 220 Asn Asp Gln Cys Lys Gly Phe Phe Pro Pro Pro Val Pro Gly Ser Ser 225 235 240 Arg Ser Tyr Tyr Ile Tyr Asp Asp Met Val Cys Ala Ala Asp Tyr Asp 255 255 Met Ser Lys Ser Ile Cys Ala Gly Asp Ser Gly Gly Pro Leu Val Cys  $260 \hspace{1cm} 265 \hspace{1cm} 270 \hspace{1cm}$ Leu Leu Glu Gly Ser Trp Tyr Val Val Gly Leu Thr Ser Trp Ser Ser 275 280 285 Thr Cys Glu Glu Pro Ile Val Ser Pro Ser Val Phe Ala Arg Val Ser 290 295 300 Tyr Phe Asp Lys Trp Ile Lys Asp Asn Lys Lys Ser Ser Ser Asn Ser 305 310 315 320 Lys Pro Gly Glu Ser Pro His His Pro Gly Ser Pro Glu Asn Glu Asn 325 330 335 Pro Glu Gly Asn Asn Lys Asn Gln Gly Thr Val Ile Lys Pro Val Cys 340 345 350Page 141

Thr Ala Leu Leu Ser Gln Thr Leu Leu Gln Gln Leu Ile 355 360 365

<210> 62 <211> 143 <212> PRT

<213> Mus musculus

Met Ser Val Gln Lys Val Ile Val His Lys Asp Tyr Asn Arg Phe His 30 Pro Glu Glu Asn Ile Lys Ser Arg Thr Gln Gly Ser Asp Ile Val Leu Leu Gln Leu Arg Ser Ser Val Glu Lys Ala Cys Val Pro Glu Glu Asn Ile Lys Ser Ser His Ile Leu Pro Ala Ser Gly Trp Gly Tyr Leu Arg Ser Ser Ser Ser Mis Ile Lys Ala Cys Trp Ala Ser Gly Trp Gly Tyr Leu Arg Ser Ser Mis Ile Lys Ala Cys Trp Ala Ser Gly Trp Gly Tyr Leu Arg 80 Glu Asp Val Arg Ile Pro Leu Pro Asn Glu Leu Tyr Glu Ala Glu Leu Pro Gly Ser Gly Arg Ser Tyr Tyr Ile Tyr Asp Asp Met Ser Lys Ser Ile Cys Ala Gly Leu Leu Leu Leu Leu Leu Cys Ala Asp Tyr Asp Met Ser Lys Ser Ile Cys Ala Gly Leu Leu Leu Leu Leu Leu Cys Ala Arg Ile Cys Ala Gly Leu Leu Leu Leu Leu Cys Ala Arg Ile Cys Ala Gly Leu Leu Leu Leu Leu Leu Cys Ala Asp Tyr Asp Met Ser Lys Ser Ile Cys Ala Gly Leu Leu Leu Leu Leu Cys Ala Asp Tyr Asp Met Ser Lys Ser Ile Cys Ala Gly Leu Leu Leu Leu Cys Ala Gly Leu Leu Leu Cys Ala Gly Leu Leu Leu Cys Ala Gly Cys Ala Gly Leu Leu Leu Cys Ala Gly Leu Leu Leu Cys Ala Gly Leu Cys Ala Gly Leu Leu Leu Cys Ala Gly Cys Ala Gly

<210> 63 <211> 273 <212> PRT <213> Ovis aries

<400> 63
Met Leu His Leu Leu Ala Leu Ala Leu Leu Leu Ser Leu Val Ser Ala
Ala Pro Ala Pro Gly Gln Ala Leu Gln Arg Ser Gly Ile Ile Gly Gly
Lys Glu Ala Pro Gly Ser Arg Trp Pro Trp Gln Val Ser Leu Arg Val
Arg Asp Gln Tyr Trp Arg His Gln Cys Gly Gly Ser Leu Ile His Pro
50
Gln Trp Val Leu Thr Ala Ala His Cys Ile Gly Pro Glu Leu Gln Glu
65
Pro Ser Asp Phe Arg Val Gln Leu Arg Glu Gln His Leu Tyr Tyr Gln
80

Asp Arg Leu Leu Pro Ile Ser Arg Val CURA2221.APP | His Tyr Tyr Met Val Glu Asn Gly Ala Asp Ile Ala Leu Leu Gln Leu Glu Glu Glu Pro 130 | Thr 140 |

<210> 64 <211> 273 <212> PRT

<213> Ovis aries

Met Val Glu Asn Gly Ala Asp Ile Ala Leu Leu Gln Leu Glu Glu Pro Val Ser Ile Ser Cys His Val Arg Pro Val Thr Leu Pro Pro Ala Ser Glu Thr Phe Pro Pro Gly Ser Gln Cys Trp Val Thr Gly Trp Gly Asn 145 Thr Phe Pro Pro Gly Ser Gln Cys Trp Val Thr Gly Trp Gly Asn 160 Val Asp Asn Gly Arg Pro Leu Pro Pro Pro Pro Tyr Pro Leu Lys Gln Val 175 Val Lys Val Pro Ile Val Glu Asn Ser Val Cys Asp Trp Lys Tyr His Ser Gly Leu Ser Thr Asp Tyr Ser Val Pro Ile Val Gln Glu Asp Asn Leu 195 Thr Asp Gly Arg Asp Ser Cys Gln Gly Asp Ser Gly Gly Pro Leu Val Cys Lys Val Asn Gly Thr Trp Leu Gln Ala Gly Val Val 240 Ser Trp Gly Asp Gly Cys Ala Asn Pro Asp Tyr Pro Gly Val Tyr Thr Arg Ile Thr Ser Tyr Leu Asp Trp Ile His Gln Tyr Val Pro Gln Glu Pro Pro

<210> 65 <211> 205 <212> PRT

<213> Artificial Sequence

<400> 65
Ser Ala Pro Ala Ser Ser Val Arg Val Ser Leu Ser Val Arg Leu Gly
Glu His Asn Leu Ser Leu Thr Glu Gly Thr Glu Gln Lys Phe Asp Val
Lys Lys Thr Ile Ile Val His Pro Asn Tyr Asn Pro Asp Thr Leu Asp
Asn Gly Ala Tyr Asp Asn Asp Ile Ala Leu Leu Lys Leu Lys Ser Pro
Gly Val Thr Leu Gly Asp Thr Val Arg Pro Ile Cys Leu Pro Ser Ala
80
Ser Ser Asp Leu Pro Val Gly Thr Thr Cys Thr Val Ser Gly Trp Gly
95

Arg Arg Pro Thr Lys Asn Leu Gly Leu CURA2221.APP Ser Asp Thr Leu Gln Glu Val Val Val Pro 110 Val Ser Arg Glu Thr Cys Arg Ser Ala Tyr Glu Tyr Gly Gly Thr Asp Asp Lys Val Glu Phe Val Thr Asp Asn Met Ile Cys Ala Gly Ala Leu Gly Gly Lys Asp Ala Cys Gln Gly Asp Ser Gly Gly 160 Pro Leu Val Cys Ser Asp Gly Asn Arg Asp Gly Arg Trp Glu Leu Val Gly Ile Val Ser Tyr Gly Ser Tyr Gly Cys Ala Arg Gly Asn Lys Pro Gly Val Tyr Thr Arg Val Ser Ser Tyr Leu Asp Trp Ile 205

<210> 66 <211> 349 <212> PRT

<213> Homo sapiens

 <400> 66 Met Asn Arg
 Lys
 Ala Leu Arg
 Cys
 Leu Gly His Leu Phe Leu Ser Leu 15

 Gly Met Val
 Cys
 Leu Arg
 Ile Gly Gly Phe Ser Ser Val Val Ala Ala Leu 30
 Ala Leu 30

 Gly Ala Thr Ile Ile Cys
 Asn Lys Ile Pro Gly Leu Ala Pro Arg Gln Arg Ala Ile Ile Val Ile Gly Glu 60
 Arg Ala Ile Cys Gln Ser Arg Pro Asp Ala Ile Ile Val Ile Gly Glu Glu 60

 Arg Ser Gln Met Gly Leu Asp Glu Cys Gln Phe Gln Phe Arg Asn Gly 75
 Asp Glu Cys Gln Phe Gln Phe Arg Asp Gly 80

 Arg Trp Asn Cys Ser Ala Leu Gly Glu Arg Thr Val Phe Gly Lys Glu 95

 Leu Lys Val Gly Ser Arg Asp Gly Ala Phe Thr Tyr Ala Ile Ile Ile Ala 110

 Ala Gly Val Ala His Ala Ile Thr Ala Ala Cys Thr His Gly Asn Leu 125

 Ser Asp Cys Gly Cys Asp Lys Glu Lys Gln Gly Gln Tyr His Arg Asp 135

 Glu Gly Trp Lys Trp Gly Gly Cys Ser Ala Asp Ile Arg Tyr Gly Ile 160

 Gly Phe Ala Lys Val 165
 Phe Val Asp Ala Asp Glu Ile Lys Gln Asp Lys Ile Leu 170

 Arg Thr Leu Met Asn Leu His Asn Asn Glu Ala Gly Arg Lys Ile Leu 180

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Glu Glu Asn Met Lys Leu Glu Cys Lys Cys His Gly Val Ser Gly Ser Cys Thr 195 Thr Lys Thr Cys Trp Thr Thr Leu Pro Gln Phe Arg Glu Leu Cys Tyr Val Leu Lys Asp Lys Tyr Asn Glu Ala Val His Val Glu Pro 240 Val Arg Ala Ser Arg Asn Lys Arg Pro Thr Phe Leu Lys Ile Lys Lys Cys Ser Pro Asn Tyr Cys Glu Glu Asp Pro Val Tyr Asp Leu Val Tyr Ile Glu Cys Ser Pro Asn Tyr Cys Glu Glu Asp Pro Val Thr Gly Ser Val Gly Cys Asp Leu Met Cys Ash Cys Asp Cys Tyr Asn Thr Gln Cys Ash Cys Lys Phe His Typ Cys Cys Tyr Val Lys Cys Asn Thr Cys Ser Glu Arg Thr Glu Met Tyr Thr Cys Lys Cys Asn Thr Cys Ser Glu Arg Thr Glu Met Tyr Thr Cys Lys Cys Asn Thr Cys Ser Glu Arg Thr Glu Met Tyr Thr Cys Lys Lys Lys Cys Asn Thr Cys Ser Glu Arg Thr Glu Met Tyr Thr Cys Lys

<210> 67 <211> 349 <212> PRT

<213> Homo sapiens

 Glu Glu Glu Trp Lys Trp Lys Trp 150
 Glu Glu Cys Ser Ala Asp 11e Arg Tyr Glu 11e 160

 Glu Phe Ala Lys Val 165
 Phe Val Asp Ala Asp 170
 Glu Ile Lys Glu Asp Ala 175

 Arg Thr Leu Met 180
 Asn Leu His Asn Asn 185
 Glu Ala Glu Arg Lys Ile Leu 190

 Glu Glu Asn Met Lys Leu Glu Cys Lys Cys His Glu Val Ser Glu Ser 205
 Ser Glu Ser 205

 Cys Thr Thr Lys Thr Cys Trp Thr Thr Leu Pro 200
 Phe Arg Glu Leu 200

 Gly Tyr Val Leu Lys Asp Lys Tyr Asn Glu Ala Val His Val Glu Pro 240

 Val Arg Ala Ser Arg Asn Lys Arg Pro 255
 Phe Leu Lys Lys Lys Lys 190

 Pro Leu Ser Tyr Arg Lys Pro Met Asp Thr Asp Leu Val Tyr Ile Glu 280

 Lys Ser Pro Asn Tyr Cys Glu Glu Asp Pro Val Thr Gly Ser Val Gly 280

 Thr Gln Gly Arg Ala Cys Asp Lys Phe His Tyr Asn Thr Gln Ala Ser Gly Cys 325

 Asn Trp Gln Cys Asp Gly Arg Gly Tyr Asn Thr Cys Lys

135

<210> 68

<211> 349 <212> PRT

<400> 68

<213> Mus musculus

Met Thr Arg Lys Ala Arg Arg Cys Leu Gly His Leu Phe Leu Ser Leu 10 15 
Gly Ile Val Tyr Leu Arg Ile Gly Gly Phe Ser Ser Val Val Ala Leu 20 
Gly Ala Ser Ile Ile Cys Asn Lys Ile Pro Gly Leu Ala Pro Arg Gln 45 
Arg Ala Ile Cys Gln Ser Arg Pro Asp Ala Ile Ile Val Ile Gly Glu 50

Gly Ser Gln Met Gly Leu Asp Glu Cys Gln Phe Gln Phe Arg Asn Gly 65 70 75 80

CURA2221.APP Arg Trp Asn Cys Ser Ala Leu Gly Glu Arg Thr Val Phe Gly Lys Glu
85 90 95 Leu Lys Val Gly Ser Arg Glu Ala Ala Phe Thr Tyr Ala Ile Ile Ala 100 105 110 Ala Gly Val Ala His Ala Ile Thr Ala Ala Cys Thr Gln Gly Asn Leu 115 120 125 Ser Asp Cys Gly Cys Asp Lys Glu Lys Gln Gly Gln Tyr His Arg Asp 130 135 140 Glu Gly Trp Lys Trp Gly Gly Cys Ser Ala Asp Ile Arg Tyr Gly Ile 145 150 155 160 Gly Phe Ala Lys Val Phe Val Asp Ala Arg Glu Ile Lys Gln Asn Ala 165 170 175Arg Thr Leu Met Asn Leu His Asn Asn Glu Ala Gly Arg Lys Ile Leu 180 185 190 Glu Glu Asn Met Lys Leu Glu Cys Lys Cys His Gly Val Ser Gly Ser 195 200 205 Cys Thr Thr Lys Thr Cys Trp Thr Thr Leu Pro Gln Phe Arg Glu Leu 210 215 220 Gly Tyr Val Leu Lys Asp Lys Tyr Asn Glu Ala Val His Val Glu Pro 225 230 235 240 Val Arg Ala Ser Arg Asn Lys Arg Pro Thr Phe Leu Lys Ile Lys Lys 245 250 255 Pro Leu Ser Tyr Arg Lys Pro Met Asp Thr Asp Leu Val Tyr Ile Glu 260 265 270 Lys Ser Pro Asn Tyr Cys Glu Glu Asp Pro Val Thr Gly Ser Val Gly 275 280 285 Thr Gln Gly Arg Ala Cys Asn Lys Thr Ala Pro Gln Ala Ser Gly Cys 290 295 300 Asp Leu Met Cys Cys Gly Arg Gly Tyr Asn Thr His Gln Tyr Ala Arg 305 310 315 320 Val Trp Gln Cys Asn Cys Lys Phe His Trp Cys Cys Tyr Val Lys Cys 325 330 335 Asn Thr Cys Ser Glu Arg Thr Glu Met Tyr Thr Cys Lys 340 345

<sup>&</sup>lt;210> 69 <211> 349 <212> PRT

<sup>&</sup>lt;213> Mus musculus

<sup>&</sup>lt;400> 69 Met Thr Arg Lys Ala Arg Arg Cys Leu Gly His Leu Phe Leu Ser Leu 1 10 15 Gly Ile Val Tyr Leu Arg Ile Gly Gly Phe Ser Ser Val Val Ala Leu 20 25 30 Page 148

Gly Ala Ser Ile Ile Cys Asn Lys Ile Pro Gly Leu Ala Pro Arg Gln 35 40 45 Arg Ala Ile Cys Gln Ser Arg Pro Asp Ala Ile Ile Val Ile Gly Glu
50 60 Gly Ser Gln Met Gly Leu Asp Glu Cys Gln Phe Gln Phe Arg Asn Gly
65 75 80 Arg Trp Asn Cys Ser Ala Leu Gly Glu Arg Thr Val Phe Gly Lys Glu 85 90 95 Leu Lys Val Gly Ser Arg Glu Ala Ala Phe Thr Tyr Ala Ile Ile Ala 100 105 110 Ala Gly Val Ala His Ala Ile Thr Ala Ala Cys Thr Gln Gly Asn Leu 115 120 125 Ser Asp Cys Gly Cys Asp Lys Glu Lys Gln Gly Gln Tyr His Trp Asp 130 140 Glu Gly Trp Lys Trp Gly Gly Cys Ser Ala Asp Ile Arg Tyr Gly Ile 145 150 155 160 Gly Phe Ala Lys Val Phe Val Asp Ala Arg Glu Ile Lys Gln Asn Ala 165 170 175 Arg Thr Leu Met Asn Leu His Asn Asn Glu Ala Gly Arg Lys Ile Leu 180 185 190 Glu Glu Asn Met Lys Leu Glu Cys Lys Cys His Gly Val Ser Gly Ser 195 200 205 Cys Thr Thr Lys Thr Cys Trp Thr Thr Leu Pro Gln Phe Arg Glu Leu 210 220 Gly Tyr Val Leu Lys Asp Lys Tyr Asn Glu Ala Val His Val Glu Pro 225 230 235 240 Val Arg Ala Ser Arg Asn Lys Arg Pro Thr Phe Leu Lys Ile Lys Lys 255 255 Pro Leu Ser Tyr Arg Lys Pro Met Asp Thr Asp Leu Val Tyr Ile Glu 260 265 270 Leu Ser Pro Asn Tyr Cys Glu Glu Asp Pro Val Thr Gly Ser Val Gly 275 280 285 Thr Gln Gly Arg Ala Cys Asn Lys Thr Ala Pro Gln Ala Ser Gly Cys 290 295 300 Asp Leu Met Cys Cys Gly Arg Gly Tyr Asn Thr His Gln Tyr Ala Arg 305 310 315 320 Val Trp Gln Cys Asn Cys Lys Phe His Trp Cys Cys Tyr Val Lys Cys 325 330 335 Asn Thr Cys Ser Glu Arg Thr Glu Met Tyr Thr Cys Lys 340

<211> 349 <212> PRT

<213> Gallus gallus

<400> 70

Met Asn Arg Lys Thr Arg Arg Trp Ile Phe His Ile Phe Leu Ser Leu
1 5 10

Gly Ile Val Tyr Ile Lys Ile Gly Gly Phe Ser Ser Val Val Ala Leu 20 25 30

Gly Ala Ser Ile Ile Cys Asn Lys Ile Pro Gly Leu Ala Pro Arg Gln 35 40 45

Arg Ala Ile Cys Gln Ser Arg Pro Asp Ala Ile Ile Val Ile Gly Glu
50 55 60

Gly Ser Gln Met Gly Ile Asn Glu Cys Gln Phe Gln Phe Arg Asn Gly
65 70 75 80

Arg Trp Asn Cys Ser Ala Leu Gly Glu Arg Thr Val Phe Gly Lys Glu
85 90 95

Leu Lys Val Gly Ser Arg Glu Ala Ala Phe Thr Tyr Ala Ile Ile Ala 100 105 110

Ala Gly Val Ala His Ala Ile Thr Ala Ala Cys Thr Gln Gly Asn Leu 115 120 125

Ser Asp Cys Gly Cys Asp Lys Glu Lys Gln Gly Gln Tyr His Lys Glu 130 135 140

Glu Gly Trp Lys Trp Gly Gly Cys Ser Ala Asp Ile Arg Tyr Gly Ile 145 150 155 160

Gly Phe Ala Lys Val Phe Val Asp Ala Arg Glu Ile Lys Gln Asn Ala 165 170 175

Arg Thr Leu Met Asn Leu His Asn Asn Glu Ala Gly Arg Lys Ile Leu 180 185 190

Glu Glu Asn Met Lys Leu Glu Cys Lys Cys His Gly Val Ser Gly Ser 195 200 205

Cys Thr Thr Lys Thr Cys Trp Thr Thr Leu Pro Lys Phe Arg Glu Leu 210 215 220

Gly Tyr Ile Leu Lys Asp Lys Tyr Asn Glu Ala Val Gln Val Glu Pro 225 230 235 240

Val Arg Ala Ser Arg Asn Lys Arg Pro Thr Phe Leu Lys Ile Lys Lys 245 250 255

Pro Leu Ser Tyr Arg Lys Pro Met Asp Thr Asp Leu Val Tyr Ile Glu 260 265 270

Lys Ser Pro Asn Tyr Cys Glu Glu Asp Pro Val Thr Gly Ser Val Gly 275 280 285

Thr Gln Gly Arg Met Cys Asn Lys Thr Ala Gln Gln Ser Asn Gly Cys 290 295 300

Asp Leu Met Cys Cys Gly Arg Gly Tyr Asn Thr His Gln Tyr Ser Arg Page 150 Val Trp Gln Cys Asn Cys Lys Phe His Trp Cys Cys Tyr Val Lys Cys 325 330 335 Asn Thr Cys Ser Glu Arg Thr Glu Val Tyr Thr Cys Lys 340 345

<210> 71 <211> 352

<212> PRT

<213> Artificial Sequence

<223> Description of Artificial Sequence: WNT domain sequence

<400> 71 Leu Cys Arg Ser Leu Pro Gly Leu Ser Pro Arg Gln Arg Gln Leu Cys 1 5 10 15

Arg Arg Asn Pro Asp Val Met Ala Ser Val Ser Glu Gly Ala Gln Leu 20 25 30

Ala Ile Gln Glu Cys Gln His Gln Phe Arg Gly Arg Arg Trp Asn Cys 40 45

Ser Thr Leu Asp Ser Leu Asn Glu Arg Ser Val Phe Gly Lys Val Leu 50 60

Lys Lys Gly Thr Arg Glu Thr Ala Phe Val Tyr Ala Ile Ser Ser Ala 65 70 75 80

Gly Val Ala His Ala Val Thr Arg Ala Cys Ser Glu Gly Glu Leu Glu 85 90 95

Ser Cys Gly Cys Asp Asp Lys Arg Lys Ala Asp Glu Glu Arg Leu Arg 100 105 110

Ile Lys Leu Glu Pro Lys Gly Pro Gly Gly Pro Gln Gly Ser Trp Lys 115 120 125

Trp Gly Gly Cys Ser Asp Asn Val Glu Phe Gly Ile Arg Phe Ser Arg 130 135 140

Glu Phe Val Asp Ala Arg Glu Arg Glu Lys Leu Met Thr Lys Ser Arg 145 150 155 160

Asp Arg Asp Ala Arg Ser Leu Met Asn Leu His Asn Asn Glu Ala Gly
165 170 175

Arg Lys Ala Val Lys Ser His Met Arg Arg Glu Cys Lys Cys His Gly 180 185 190

Val Ser Gly Ser Cys Ser Leu Lys Thr Cys Trp Leu Ser Leu Pro Asp 195 200 205

Phe Arg Glu Val Gly Asp Leu Leu Lys Glu Lys Tyr Asp Gly Ala Ile 210 215 220

Glu Val Glu Val Asn Lys Arg Gly Lys Gly Gln Arg Ser Leu Ser Ser 225 230 235 240 Page 151

Arg Lys Gln Ala Ser Ala Leu Glu Ala Ala Asn Glu Arg Phe Lys Lys 245 250 255 Pro Thr Arg Asn Gln Tyr Thr Asp Leu Val Tyr Leu Glu Lys Ser Pro 260 265 270 Asp Tyr Cys Glu Arg Asp Arg Glu Thr Gly Ser Leu Gly Thr Gln Gly 275 280 285 Arg Val Cys Asn Lys Thr Ser Lys Gly Leu Gln Trp Arg Asp Gly Cys 290 295 300 Glu Leu Leu Cys Cys Gly Arg Gly Tyr Asn Thr Glu Gln Lys Val Glu 305 310 315 320 Arg Thr Glu Lys Cys Asn Cys Lys Phe His Asn Gly Trp Cys Cys Tyr 325 330 335 Val Lys Cys Glu Glu Cys Thr Glu Val Val Glu Val His Thr Cys Lys 340 350

<210> 72 <211> 1216 <212> PRT

<213> Rattus norvegicus

<400> 72 Met Cys Leu Pro Ser Cys Leu Leu Ser Ile Trp Val Leu Phe Met Ala 1 5 10 15 Ala Gln Ser Leu Gly Lys Thr Trp Val Pro Asp His Cys Arg Ser Pro
20 25 30 Thr Glu Ala Thr Cys Asn Phe Val Cys Asp Cys Gly Asp Cys Ser Asp 40 45 Glu Ala Gln Cys Gly Phe His Gly Ala Ser Thr Thr Pro Asn Thr Pro 50 55 60 Phe Thr Cys Asn Phe Glu Gln Asp Pro Cys Gly Trp Gln Asp Ile Ser 65 70 75 80 Thr Ser Gly Tyr Arg Trp Leu Arg Asp Arg Ala Gly Ala Gly Leu Asp
85
90
95 Ser Ser Gly Pro His Ser Asp His Thr Arg Gly Thr Asp Leu Gly Trp 100 105 110 Tyr Met Ala Val Gly Thr His Ser Gly Lys Glu Pro Ser Thr Arg Thr 115 120 125 Leu Arg Ser Pro Val Met Arg Glu Ala Ala Pro Thr Cys Glu Leu Arg 130 135 140 Leu Trp Tyr His Thr Asp Ser Arg Asp Val Ala Glu Leu Arg Leu Asp 145 150 155 160 Leu Thr His Gly Met Glu Thr Leu Thr Leu Trp Gln Ser Ser Gly Pro Page 152

Trp Gly Pro Trp Pro Gly Arg Glu Leu Ala Val Asn Thr Gly Arg Ile 180 185 190 Gln Gly Asp Phe Lys Val Thr Phe Ser Ala Thr Arg Asn Ala Thr His 195 200 205 Gly Ala Val Ala Leu Asp Asp Met Glu Phe Trp Asp Cys Gly Leu 210 215 220 Pro Ile Pro Gln Ala Arg Cys Pro Leu Gly His His His Cys Gln Asn 235 240 Lys Ala Cys Val Glu Pro His Gln Leu Cys Asp Gly Glu Asp Asn Cys 245 250 255 Gly Asp Ser Ser Asp Glu Asp Pro Leu Ile Cys Ser His His Met Ala 260 265 270 Thr Asp Phe Glu Thr Gly Leu Gly Pro Trp Thr Gln Leu Glu Gly Trp 275 280 285 Thr Arg Asn Phe Ser Ala Gly Ser Met Val Ser Pro Ala Trp Pro His 290 295 300 Arg Asp His Ser Arg Asn Ser Ala Tyr Gly Phe Phe Leu Val Ser Val 305 310 315 320 Ala Lys Pro Gly Thr Thr Ala Val Leu Tyr Ser Pro Glu Phe Gln Gly 325 330 335 Ser Val Ser Tyr Asn Cys Ser Phe Thr Phe Tyr Tyr Leu His Gly 340 345 350 Ser Glu Ala Asn Gln Phe Gln Leu Phe Val Gln Ala Gln Gly Leu Asn 355 360 365 Thr Thr Gln Pro Pro Val Leu Leu Arg Ser Arg His Gly Glu Leu Gly 370 375 380 Thr Ala Trp Val Arg Asp Arg Val Asn Ile Gln Ser Ala His Pro Phe 385 390 395 400 Arg Ile Leu Leu Ala Gly Glu Thr Gly Pro Gly Gly Phe Val Gly Leu 405 410 415 Asp Asp Leu Ile Met Ser Asn His Cys Ile Leu Val Pro Gly Met Ser 420 425 430 Thr Leu Gln Ser Ser Leu Ser Gly Pro Val Pro Leu Ala Leu Tyr Pro 435 440 445 Gln Thr Ser Ile Lys Arg Thr Cys Asp Ala Gly His Leu Ser Cys Asp 450 455 460 Glu Leu Cys Val Pro Pro Glu Gln Leu Cys Asp Phe Gln Gln His Cys 465 470 475 480 Ala Glu Gly Glu Asp Glu Glu Lys Cys Gly Thr Thr Asp Phe Glu Ser 485 490 495 Ala Ser Ala Gly Gly Trp Glu Asp Ile Ser Ile Gly Lys Leu Gln Trp Page 153

Gln Arg Ala Glu Ala Gln Glu Ser Gly Lys Pro Ala Arg Asp Thr Asn 515 520 525 Arg Asn Ala Pro Gly His Phe Leu Ser Leu Arg Lys Ala Trp Gly Gln
530 540 Leu Arg Ser Glu Ala Arg Ala Leu Thr Pro Thr Leu Gly Pro Ser Gly 545 550 560 Pro His Cys Glu Leu His Met Thr Tyr Tyr Phe His Ser His Pro Gln 575 Gly Phe Leu Ala Leu Ala Val Val Glu Asn Gly Phe Arg Glu Leu Leu 580 590 Trp Gln Ala Pro Ser Ser Ser Gly Gly Trp Thr Leu Gln Lys Ile 595 600 605 Leu Leu Gly Ala Arg Arg Trp Pro Phe Gln Leu Glu Phe Val Ser Leu 610 620 Val Asp Leu Asp Gly Pro Gly Gln Gln Gly Ala Gly Val Asp Asn Val 625 630 635 640 Thr Leu Arg Asp Cys Asn Pro Met Val Thr Thr Glu Ser Asp Gln Glu 645 650 655 Val Ser Cys Asn Phe Glu Arg Asp Ser Cys Ser Trp His Thr Gly His 660 665 670 Leu Thr Asp Ala His Trp His Arg Val Lys Ser His Gly Ser Gln Tyr 675 680 685 Asp His Thr Thr Gly Gln Gly Phe Phe Met Phe Leu Asp Pro Met Asp 690 695 700 Pro Pro Ala Arg Gly Gln Gly Ala Leu Leu Leu Thr Arg Pro Gln Val 705 710 715 720 Pro Val Val Pro Lys Glu Cys Leu Ser Phe Trp Tyr His Leu His Gly
725 730 735 Pro Gln Ile Gly Thr Leu Cys Leu Ala Met Arg Arg Glu Glu Glu 740 745 750 Asp Thr Leu Leu Trp Ser Arg Ser Gly Thr His Gly Asn Arg Trp His 755 760 765 Gln Ala Trp Val Thr Leu His His Gln Leu Gln Pro Ser Thr Lys Tyr 770 775 780 Gln Leu Leu Phe Glu Gly Leu Arg Asp Gly Tyr His Gly Thr Met Gly 785 790 795 800 Leu Asp Asp Met Ala Val Arg Pro Gly Pro Cys Trp Ala Ala Lys Arg 805 810 815 Cys Ser Phe Glu Asp Ser Asp Cys Gly Phe Ser Pro Gly Asp Trp Gly 820 825 830 Leu Trp Thr Arg Gln Asn Asn Ala Ser Gly Leu Gly Pro Trp Gly Pro Page 154

Trp Ile Asp His Thr Thr Gly Thr Ala Gln Gly His Tyr Met Val Val 850 860 Asp Thr Ser Pro Asn Leu Leu Pro Lys Gly His Val Ala Ser Leu Thr 865 870 875 880 Ser Glu Glu His Pro Pro Leu Ser Arg Pro Ala Cys Leu Ser Phe Trp 885 890 895 Tyr His Leu Ser Phe His Asn Pro Gly Thr Leu Arg Val Phe Val Glu 900 905 910 Glu Ser Thr Arg Arg Gln Glu Leu Ser Ile Ser Gly His Gly Gly Phe 915 920 925 Ala Trp Arg Leu Gly Ser Val Asn Val Gln Ala Glu Gln Ala Trp Lys 930 935 940 Val Val Phe Glu Ala Met Ala Ser Gly Val Glu His Ser Tyr Met Ala 945 950 955 960 945 Leu Asp Asp Ile Ser Leu Gln Asp Gly Pro Cys Ala Gln Pro Gly Ser 965 970 975 Cys Asp Phe Glu Ser Gly Leu Cys Gly Trp Ser His Leu Pro Trp Pro 980 985 990 Gly Leu Gly Gly Tyr Ser Trp Asp Trp Ser Ser Gly Ala Thr Pro Ser 995 1000 1005 Arg Tyr Pro Arg Pro Ser Val Asp His Thr Val Gly Thr Glu Ala Gly 1010 1020 His Phe Ala Phe Phe Glu Thr Ser Val Leu Gly Pro Gly Gln Ala 1025 1030 1035 1040 Ala Trp Leu Gly Ser Glu Pro Leu Pro Ala Thr Ala Val Ser Cys Leu 1045 1050 1055 His Phe Trp Tyr Tyr Met Gly Phe Pro Ala His Phe Tyr Lys Gly Glu 1060 1065 1070 Leu Arg Val Leu Leu Ser Ser Thr Gln Gly Gln Leu Ala Val Trp His 1075 1080 1085 Arg Gly Gly His Leu Arg Asp Gln Trp Leu Gln Val Gln Ile Glu Val 1090 1095 1100 Ser Ser Ser Glu Glu Phe Gln Ile Val Phe Glu Ala Thr Leu Gly Gly 1110 1115 Gln Pro Ala Leu Gly Pro Ile Ala Leu Asp Asp Val Glu Tyr Leu Ala 1125 1130 1135

Gly Gln His Cys Lys Gln Pro Thr Pro Ser Gln Gly Arg Val Ala Ala 1140 1145 1150

Pro Val Ser Val Pro Val Ala Val Gly Gly Ala Leu Leu Leu Phe Leu 1155 1160 1165

Leu Leu Cly Leu Gly Gly Trp His Trp Leu Gln Lys Gln His Leu Page 155

Pro Cys Gln Ser Thr Asp Ala Ala Ala Ser Gly Phe Asp Asn Ile Leu 1185 1190 1195 1200

Phe Asn Ala Asp Gln Val Thr Leu Pro Glu Ser Ile Thr Ser Asn Pro

<210> 73

<211> 688

<212> PRT

<213> Xenopus laevis

<400> 73

Met Met Leu Ser His Trp Val Leu Leu Leu Ser Leu Gly Ala Val Trp 1 5 10 15

Leu Ala Glu Gly Glu Ile Ser Pro Gly Ser Cys Thr Phe Glu Asn 20 25 30

Ser Thr Cys Ala Tyr Thr Ser Ala Phe Pro Phe Leu Gln Trp Thr Val $^{35}$   $^{40}$   $^{45}$ 

Asn Ile Glu Gly His Tyr Val Ser Val Asp Ser Ser Asn Gly Leu Arg 50 55 60

Gly Gln Lys Ala Val Leu Ile Ser Pro Asp Leu His Leu Ala Glu Trp 65 70 75 80

Ser Cys Leu Arg Leu Val Tyr Gln Ile Ala Gly Ser Glu Ser Ser Pro  $85 \hspace{1cm} 90 \hspace{1cm} 95$ 

Ser Pro Ser Ser Leu Asn Val Phe Val Arg Pro Glu Gly Glu Ser Phe 100 105 110

Asp Tyr Leu Leu Trp Ser Ala Glu Glu His Ser Asp Ser Trp Leu Ile 115 120 125

Ser Ser Ile Asp Leu Lys Asn Thr Thr Lys Arg Phe Lys Ile Ile Leu 130 140

Glu Gly Val Leu Gly Glu Asn Thr Met Ser Ser Ile Ala Ile Phe Glu 145 150 155 160

Val Lys Met Thr Thr Gly Tyr Cys Ile Glu Cys Asp Phe Glu Glu Asn 165 170 175

His Leu Cys Gly Tyr Met Asn Ser Trp Asn Pro Asn Val Asn Trp Phe 180 185 190

Val Gly Gly Asn Val Lys Asn Ser His Ser Ile Leu Pro Arg Asp 195 200 205

His Thr Leu Asn Asn Glu Leu Gly His Tyr Met Tyr Val Asp Ser Val 210 220

Tyr Val Lys His Phe Gln Glu Val Ala Gln Leu Val Ser Pro Leu Ile 225 230 235 240 CURA2221.APP

Ile Thr Pro Ile Ser Gly Cys Leu Ser Phe Tyr Tyr Gln Leu Gln Arg
245 250 255 Glu Thr Ser Asn Ile Phe Leu Val His Thr Arg Asp Leu His Gly Ser 260 265 270 Tyr Asp Glu Ile Trp Lys Met Gly Ala Val Arg Gln Gly Glu Trp Asn 275 280 285 Leu Ala Glu Val Asp Leu Asn Ala His Val Pro Leu Glu Val Ile Phe 290 295 300 Glu Val Ala Phe Asn Gly Ile Gln Ala Gly Tyr Val Ala Leu Asp Asp 305 310 315 320 Ile Leu Phe Ser Pro Val Ser Cys Ser Gly Gln Glu Gly Met Phe Phe 325 330 335 Asp Ala Arg Glu Ala Gly Cys Asp Phe Glu Glu Gly Met Cys Gln Phe 340 345 350 His Gln Asp Asp Asn Asn Gly Ser Gly Trp Ser Arg Val Lys 355 360 365 Pro Asn Ala Tyr Gln Met Gly Asp His Thr Thr Gly Leu Gly Tyr Phe 370 380 Met Ile Ala Asn Thr Arg Phe Thr Gly Gln Pro Ala Tyr Phe Gly Arg 385 390 395 400 Leu Tyr Gly Pro Ser Leu Pro Gly Asn Ile Gln Tyr Cys Ile Arg Phe  $405 \hspace{1cm} 410 \hspace{1cm} 415$ Phe Tyr Ser Leu Tyr Gly Phe Tyr Lys Thr Ile Asp Ser Leu Ala Val 420 425 430 Tyr Ile Phe Glu Glu Asn His Val Val Gln Glu Lys Ile Trp Ser Ala 435 440 445 His Glu Thr Pro Lys Gly Val Trp Leu Gln Ala Glu Ile Ser Ile His 450 455 460 Pro Met Pro Phe Lys Val Val Phe Val Ser Trp Cys Lys Ser Leu 470 475 480 Trp Asp Cys Gly Ile Ala Ala Leu Asp Asp Ile Ser Val Ser Ile Gly 485 490 495 Ser Cys Lys Ile Ser Asp Arg Ile Pro Pro Leu Pro Gly Lys Cys Thr 500 505 510 Phe Glu Lys Asn Asp Cys Gly Phe Gly Ala Gly Met Ala Lys Glu Gly 515 520 525 Tyr Leu Ala Gln Asn Thr Arg Glu Asp Pro Thr Phe Tyr Thr Gly Pro 530 535 540 Asn Gly Asp His Thr Ser Gly Val Gly Tyr Tyr Met Tyr Ile Glu Ala 545 550 555 560 Thr Asn Met Val Phe Gly Gln Lys Ala Lys Leu Ile Ser Arg Pro Leu 565 570 575

Arg Ala Val Ala Gly Lys Gln Cys Leu CURA2221.APP
Thr Phe Tyr Tyr His Met Tyr
Gly Ala Gly Thr Gly Leu Leu Asn Val Tyr Leu Thr Lys Glu Gly Asp
S95 Thr Leu Leu Asn Val Tyr Leu Thr Lys Glu Gly Asp
1le Asn Lys Asp Thr Leu Leu Trp Thr Arg Lys Gly Glu Gln Ser Ile
Thr Trp Leu Lys Ala Gln Met Glu Tyr Glu Ser Glu Gln Gln His Lys
625 The Val Phe Glu Ala Val Arg Gly Ile Ser Ile Arg Ser Asp Ile Ala
655 Ala
1le Asp Asp Ile Leu Phe Gln Asn Gly Pro Cys Asn Asp Ser Ser Asp
Pro Leu Gln Ser Ser Gly Tyr Ser Asp Asn Phe Asn Asn Ile Glu Phe

<210> 74 <211> 5376 <212> PRT

<213> Mus musculus

<400> 74
Met Ala Leu Pro Val Trp Thr Leu Met Leu Leu Val Gly Ala Ala Trp
Gly Gln Glu Gln Val Pro Ala Trp Arg Pro Asn Ser Pro Asp Leu Gly
Pro Met Val His Thr Ser Arg Glu Asp Ser Ile Leu Ser Lys Cys Asp
Phe Glu Asp Asn Ser Arg Pro Phe Cys Asp Trp Ser Gln Met Ser Ala
Asp Asp Gly Asp Trp Ile Arg Thr Thr Gly Pro Ser Leu Thr Gly Thr
65 Asp Pro Pro Gly Gly Tyr Pro Asn Gly Glu Gly Tyr Tyr Leu His
Met Asp Pro Lys Thr Phe Pro Gln Gly Gly Val Ala Arg Leu Arg Ser
Pro Asp Ile Trp Glu Gln Gly Pro Leu Cys Val His Phe Ala Phe His
Met Phe Gly Leu Ser Trp Gly Ala Gln Leu Arg Leu Leu Leu Arg
Gly Arg Lys His Leu Arg Pro Tyr Val Leu Trp Lys His Val Asn Thr
160
Gln Ser Pro Ser Trp Met Pro Thr Thr Val Tro Ala Asp His
Pro Asp Ile Tro Ala Asp His
Pro Asp Ile Tro Ser Trp Met Pro Thr Thr Val Tro Ala Asp His
Pro Asp Lys His Leu Arg Pro Tyr Val Leu Trp Lys His Val Asn Thr
160
Page 158

Asp Ile Pro Ser Trp Leu Met Phe Glu Gly Met Arg Gly Asn Thr Ala 180 185 190 Tyr Leu Asp Ile Ser Leu Asp Gly Leu Ser Ile Gln Arg Gly Thr Cys 195 200 205 Asn Gln Val Cys Met Ser Gln Met Cys Thr Phe Asp Thr Leu Asn Asp 210 215 220 Leu Cys Gly Trp Ser Trp Val Pro Thr Ala Thr Gly Ala Lys Trp Thr 225 230 235 240 Gln Lys Lys Gly Pro Thr Gly Lys Gln Gly Val Gly Pro Ala Glu Asp 245 250 255 Phe Ser Asn Pro Gly Asn Gly Tyr Tyr Met Leu Leu Asp Ser Thr Asn 260 265 270 . Ala Arg Pro Gly Gln Lys Ala Val Leu Leu Ser Pro Leu Ser His Ser 275 280 285 Arg Gly Cys Met Thr Leu Ser Phe His Tyr Ile Met His Gly Gln Gly 290 295 300His Glu Glu Gly Leu Phe Val Tyr Ala Thr Phe Leu Gly Asn Ile Arg 305 310 315 320 Lys Tyr Thr Leu Phe Ser Gly His Pro Gly Pro Asp Trp Gln Ala Val 325 330 335 Ser Val Asn Tyr Thr Gly Gln Gly Gln Ile Gln Phe Met Val Val Gly 340 345 350 Met Phe Gly Asn Ile Pro Glu Pro Ala Ile Ala Val Asp Ala Ile Ser 355 360 365 Ile Ala Pro Cys Gly Glu Ser Phe Pro Gln Cys Asp Phe Glu Asp Arg 370 375 380 Val His Pro Phe Cys Asp Trp Asn Gln Val Tyr Gly Asp Met Gly His 385 390 395 400 385 Trp Ser Trp Gly Ser Lys Ser Val Pro Thr Leu Ile Ala Gly Ser Pro 405 410 415 Arg Glu Phe Pro Tyr Gly Gly Glu His Tyr Ile Phe Phe Asp Ser Val 420 425 430 Lys Leu Ser Gln Glu Gly Gln Ser Ala Arg Leu Val Ser Pro Pro Phe 435 440 445 Cys Ala Pro Gly Gly Ile Cys Val Glu Phe Ala Tyr His Met Tyr Gly
450 460 Leu Gly Lys Gly Thr Thr Leu Lys Leu Leu Gly Ser Pro Ala Gly 465 470 475 480 Ser Ser Pro Ile Pro Leu Trp Asn Arg Val Gly Ser Gln Ser Ser Gly 485 490 495 Trp Met Asn Ser Ser Val Thr Ile Pro Lys Gly Tyr Gln Gln Pro Met 505 Page 159

Gln Leu Phe Ile Glu Ala Thr Arg Gly Thr Ser Thr Ala Phe Val Val 515 520 525 Ala Leu Asn Phe Ile Leu Ile Ser His Gly Pro Cys Arg Val Leu Leu 530 540 Gln Thr Glu Ile Pro Ser Ser Pro Leu Leu Pro Pro Thr Gly Pro Ser 545 550 555 560 Glu Ser Thr Val Pro Thr Leu Pro Met Glu Gln Pro Thr Ser Pro Thr 565 570 575 Lys Ala Thr Thr Val Thr Ile Glu Ile Pro Thr Thr Pro Thr Glu Glu 580 590 Ala Thr Ile Pro Thr Glu Thr Thr Thr Val Pro Thr Glu Val Ile Asn 595 600 605 Ser Pro Lys Glu Thr Ser Ile Pro Pro Glu Val Thr Ile Pro Thr 610 620 Glu Val Ile Thr Val Ser Pro Glu Glu Ile Ile Ser Pro Thr Glu Val 625 630 635 640 Thr Pro Val Pro Thr Asp Val Thr Ala Ala Tyr Val Glu Ala Thr Asn 645 650 655 Ala Ser Pro Glu Glu Thr Ser Val Pro Pro Glu Val Thr Ile Leu Thr 660 665 670 Glu Val Thr Thr Val Ser Pro Glu Glu Thr Thr Val Pro Thr Glu Val 675 680 685 Pro Ile Val Leu Ile Glu Ala Thr Ala Phe Pro Thr Gly Glu Thr Thr 690 695 700 Leu Tyr Thr Glu Val Pro Thr Val Pro Thr Glu Val Thr Gly Val His 705 710 715 720 Thr Glu Val Thr Asn Val Ser Pro Glu Glu Thr Ser Val Pro Thr Glu
725 730 735 Glu Thr Ile Ser Thr Glu Val Thr Thr Val Ser Pro Glu Glu Thr Thr 740 745 750 Val Pro Thr Glu Val Pro Ile Val Leu Ile Glu Ala Thr Ala Ser Pro 755 760 765 Thr Gly Glu Ile Thr Leu Tyr Thr Glu Val Pro Thr Val Pro Thr Glu 770 775 780 Val Thr Gly Val His Thr Glu Val Thr Asn Val Ser Pro Glu Glu Thr 785 790 795 800 Ser Val Pro Thr Glu Glu Thr Ile Ser Thr Glu Val Thr Thr Val Ser 805 810 815 Pro Glu Glu Thr Thr Leu Pro Thr Glu Val Pro Thr Val Ser Thr Glu 820 825 830 Val Thr Asn Val Ser Pro Glu Glu Thr Ser Val Pro Pro Glu Glu Thr 835 840 845 Page 160

Ile	Leu 850	Thr	Thr	Leu	Tyr	Thr 855	Glu	۷a٦	Pro	Thr	Val 860	Pro	Thr	Glu	٧a٦
Thr 865	Gly	va1	His	Thr	G]u 870		Thr	Asn	(Val	Ser 875		Glu	Glu	Thr	Ser 880
٧a٦	Pro	Thr	Glu	G1u 885	Thr	Ile	Ser	Thr	Glu 890		Thr	Thr	٧a٦	Ser 895	Pro
Glu	Glu	Thr	Thr 900		Pro	Thr	Glu	va1 905		Thr	۷al	Ser	Thr 910	Glu	٧a٦
Thr	Asn	Val 915	Ser	Pro	Glu	Glu	Thr 920	Ser	Val	Pro	Pro	G1u 925	Glu	Thr	Ile
Leu	Thr 930	Glu	Ile	Thr	Thr	Val 935	Ser	Pro	Glu	Glu	Thr 940	٧a٦	Phe	Pro	Ile
G]u 945	Gly	Thr	Thr	Leu	Pro 950	Thr	Glu	٧a٦	Leu	Thr 955	۷al	Pro	Ile	Glu	val 960
Thr	Thr	Phe	Pro	Thr 965	Gly	Glu	Thr	Thr	Val 970	Pro	Thr	Glu	٧a٦	Pro 975	Thr
Val	Ser	Thr	G1u 980	Met	Thr	Gly	٧al	His 985	Thr	Glu	٧al	Thr	Thr 990	val	Phe
Pro	Glu	Glu 995	Thr	Ser	Ile		Thr LOOO	Glu	٧a٦	Аlа		Va 1 L005	Leu	Pro	Ala
Ser 1	Ile LO10	Pro	Pro	Glu		Thr 1015	Thr	Thr	Pro		Glu 1020	val	Thr	Thr	Thr
Pro 1025	Pro	Glu	Glu	Thr	Thr L030	Ile	Pro	Ala		Va 1 1035	Thr	Thr	val	Pro 1	Pro .040
Ala	Ser	Ile	Pro	Pro 1045	Glu	Glu	Thr		Ser 1050	Leu	Thr	Glu		Thr 1055	Thr
Thr	Pro	Pro	Glu 1060	Glu	Thr	Thr	Thr 1	Pro .065	Thr	Glu	val		Thr 1070	val	Pro
Pro	Glu -	Lys L075	Thr	Thr	Ile		Thr .080	Glu	Val	Thr		Va 1 .085	Pro	Pro	Ala
Ser 1	11e .090	Phe	Pro	Glu		Thr .095	Thr	٧a٦	Pro		Glu 100	Glu	Thr	Thr	Ile
Ala 1105	Ser	Glu	Glu	Thr 1	Thr .110	Val	Ser	Thr		G]u .115	Thr	Thr	Leu	Leu 1	Thr 120
Glu	Gln	Ser	Ala 1	Val 1125	Thr	Gln	Thr	Ser 1	Ile 130	Ala	Cys	Arg		Pro 135	Cys
Pro	Ser	Pro 1	Pro L140	Leu	Met	Pro	Ile 1	Gly 145	Pro	Leu	Leu		Lys 150	Pro	Pro
Gly	val 1	Ser 155	Met	Phe	Ser	Leu 1	Ala 160	Pro	Thr	Thr	Gly 1	val 165	Ser	Thr	Thr
31u 1	Ser 170	Cys	Pro	Pro	Asn 1	Ala 175	His	Ile			180	Ala	Cys	Pro /	Ala

- Ser Cys Glu Ser Pro Lys Pro Ser Cys Gln Pro Pro Cys Ile Pro Gly 1185 1190 1195 1200

  Cys Val Cys Asn Pro Gly Phe Leu Phe Ser Asn Asn Gln Cys Ile Asn
- Cys Val Cys Asn Pro Gly Phe Leu Phe Ser Asn Asn Gln Cys Ile Asn 1205 1210 1215
- Glu Ser Ser Cys Asn Cys Pro Tyr Asn Asn Lys His Tyr Lys Pro Gly 1220 1225 1230
- Glu Glu Trp Phe Thr Pro Asn Cys Thr Glu Arg Cys Arg Cys Leu Pro 1235 1240 1245
- Gly Ser Leu Met Glu Cys Gln Ile Ser Gln Cys Gly Thr His Thr Val 1250 1255 1260
- Cys Gln Leu Lys Ser Asp Gln Tyr Gln Cys Glu Pro Tyr Gly Lys Ala 1265 1270 1275 1280
- Thr Cys Leu Val Tyr Gly Asp Leu His Phe Val Thr Phe Asp Glu Arg 1285 1290 1295
- His Ile Gly Phe Thr Gly Thr Cys Thr Tyr Ile Leu Thr Gln Thr Cys 1300 1305 1310
- Ser Asn Ser Thr Asp His Phe Phe Arg Ile Thr Ala Asn Thr Glu Glu 1315 1320 1325
- Arg Gly Val Glu Gly Val Ser Cys Leu Asp Lys Val Val Ile Ser Leu 1330 1340
- Pro Glu Thr Thr Val Thr Met Ile Ser Gly Arg His Thr Leu Ile Gly 1345 1350 1360
- Asp Gln Glu Val Thr Leu Pro Ala Ile Leu Ser Asp Asp Thr Tyr Val 1365 1370 1375
- Gly Leu Ser Gly Arg Phe Val Glu Leu Arg Thr Thr Phe Gly Leu Arg 1380 1385 1390
- Val Arg Trp Asp Gly Asp Gln Gln Leu Phe Val Thr Val Ser Ser Thr 1395 1400 1405
- Phe Ser Gly Lys Leu Cys Gly Phe Cys Gly Asn Tyr Asp Gly Asp Ser 1410 1415 1420
- Glu Leu Arg Leu Ser Trp Gln Val Glu Glu Asp Glu Asp Lys Asp Trp 1445 1450 1455
- Val Ser Ser Arg Cys Gln Lys Lys Lys Asn Pro Pro Ser Cys Asp Ala 1460 1465 1470
- Ala Leu Gly Ser Thr Met Ser Gly Pro Lys Leu Cys Gly Gln Leu Val 1475 1480 1485
- Asn Pro Ser Gly Pro Phe Glu Ala Cys Leu Leu His Leu Lys Ala Ser 1490 1495 1500
- Ser Phe Leu Asp Asn Cys Val Thr Asp Met Cys Ser Phe Gln Gly Leu 1505 1510 1515 1520 Page 162

G]n	Gln	Lys	Leu	Cys	Ala	Arg	Met	Ser	Ala	Met	Thr	Ala	Thr	Cys	Gln
	1525					1530					1535				

Asp Ala Gly Tyr Pro Val Lys Pro Trp Arg Glu Pro Gln Phe Cys Pro 1540 1545 1550

Leu Val Cys Pro Lys Asn Ser Arg Tyr Ser Leu Cys Ala Lys Pro Cys 1565 1560 1565

Pro Glu Thr Cys His Pro Ile Ser Thr Thr Gln His Cys Ser Asp Lys 1570 1575 1580

Cys Val Glu Gly Cys Glu Cys Asp Pro Gly Phe Ile Leu Ser Gly Ser 1585 1590 1595 1600

Glu Cys Val Pro Ser Ser Gln Cys Gly Cys Thr Ser Phe Gln Gly Arg 1605 1610 1615

Tyr Phe Lys Leu Gln Glu Gln Trp Phe Asn Pro Asp Cys Lys Glu Ile 1620 1625 1630

Cys Thr Cys Glu Ser His Asn His Ile Leu Cys Lys Pro Trp Lys Cys 1635 1640 1645

Lys Ala Gln Glu Ala Cys Ser Tyr Lys Asn Gly Val Leu Gly Cys His 1650 1655 1660

Ala Gln Gly Ala Ala Thr Cys Met Val Ser Gly Asp Pro His Tyr Leu 1665 1670 1675 1680

Thr Phe Asp Gly Ala Leu His His Phe Met Gly Thr Cys Thr Tyr Val 1685 1690 1695

Leu Thr Gln Pro Cys Trp Ser Lys Ser Gln Glu Asn Asn Phe Val Val 1700 1705 1710

Ser Ala Thr Asn Glu Ile His Asp Gly Asn Leu Glu Val Ser Tyr Val 1715 1720 1725

Lys Ala Val His Val Gln Val Phe Asp Leu Lys Ile Ser Met Phe Lys 1730 1735 1740

Gly Gln Lys Val Val Leu Asn Asn Gln Arg Val Val Leu Pro Val Trp 1745 1750 1760

Pro Ser Gln Gly Arg Val Thr Ile Arg Leu Ser Gly Ile Phe Val Leu 1765 1770 1775

Leu Tyr Thr Asn Phe Gly Leu Gln Val Arg Tyr Asp Gly Arg His Leu 1780 1785 1790

Val Glu Val Thr Val Pro Ser Ser Tyr Thr Gly Ser Leu Cys Gly Leu 1795 1800 1805

Cys Gly Asn Tyr Asn Asn Asn Ser Met Asp Asp Asn Leu Arg Ala Asp 1810 1815 1820

Met Lys Pro Ala Gly Asn Ser Leu Leu Leu Gly Ala Ala Trp Lys Ile 1825 1830 1835 1840

Leu Glu Ala Ser Asp Pro Gly Cys Phe Leu Ala Gly Gly Lys Pro Ser 1845 1850 1855 Page 163

- Arg Cys Ala Asp Ser Asp Met Asp Asp Val Trp Thr Lys Lys Cys Ala 1860 1865 1870
- Ile Leu Met Asn Pro Leu Gly Pro Phe Ser Asn Cys His Glu Ala Val 1875 1880 1885
- Pro Pro Gln Ala Ser Phe Ser Ser Cys Val Tyr Gly Gln Cys Glu Thr 1890 1895 1900
- Asn Gly Asp Asn Leu Thr Phe Cys His Ser Leu Gln Ala Tyr Ala Ser 1905 1910 1915 1920
- Leu Cys Ala Gln Ala Gly Gln Val Thr Trp Arg Asn Ser Thr Phe 1925 1930 1935
- Cys Pro Met Arg Cys Pro Pro Arg Ser Ser Tyr Asn Pro Cys Ala Asn 1940 1945 1950
- Ser Cys Pro Ala Thr Cys Leu Thr Leu Ser Thr Pro Arg Asp Cys Pro 1955 1960 1965
- Thr Leu Pro Cys Val Glu Gly Cys Glu Cys Gln Ser Gly His Ile Leu 1970 1975 1980
- Ser Gly Thr Thr Cys Val Pro Leu Arg Gln Cys Gly Cys Ser Asp Gln 1985 1990 1995 2000
- Asp Gly Ser Tyr His Leu Leu Gly Glu Ser Trp Tyr Thr Glu Lys Thr 2005 2010 2015
- Cys Thr Thr Leu Cys Thr Cys Ser Ala His Ser Asn Ile Thr Cys Ser 2020 2025 2030
- Pro Thr Ala Cys Lys Ala Asn His Val Cys Leu Arg Gln Glu Gly Leu 2035 2040 2045
- Leu Arg Cys Ala Ala Glu Met Gly Glu Cys Arg Ile Ser Glu Asp Ser 2050 2055 2060
- Gln Ile Val Ser Phe Asp Asp His Ser His Pro Ile Gln Asp Thr Cys 2065 2070 2075 2080
- Thr Tyr Ile Leu Val Lys Val Cys His Pro Asn Thr Asn Met Pro Phe 2085 2090 2095
- Phe Met Ile Ser Ala Lys Thr Asp Ile Asn Thr Asn Gly Lys Asn Lys 2100 2105 2110
- Thr Phe Gly Val Tyr Gln Leu Tyr Ile Asp Ile Phe Asn Phe His Ile 2115 2120 2125
- Thr Leu Gln Lys Asp His Leu Val Leu Ile Ser Leu Ile Asn Asp Ser 2130 2140
- Ile Val Thr Leu Pro Thr Thr His Ile Pro Gly Val Ser Val Met 2145 2150 2155 2160
- Thr Glu Asp Val Tyr Thr Ile Val Thr Ile Lys Asp Glu Ile Gln Val 2165 2170 2175
- Lys Phe Glu Ser Asn Asn Phe Leu Asp Val Lys Ile Pro Ala Ser Ser 2180 2185 2190 Page 164

Asn		Lys 2195	۷al	Cys	Gly		Cys 2200	Gly	Asn	Phe		G]y 2205	Glu	Glu	Glu
	G]u 210	Leu	Met	Thr		Ser 2215	Gly	Glu	Leu		G]u 2220	Asp	Glu	Gln	Glu
Phe 2225		Asn	Ser	Trp 2	Lys 2230	Asp	Lys	Ser	Met	Asp 2235	Pro	Asn	Cys	Gln	Lys 2240
Ile	Glu	Gly		Asn 2245	Leu	Gln	۷al		G]n 2250	Gln	Glu	Ile	Met	Asn 2255	Gly
Lys	Cys		Pro 2260	Ile	Asp	Phe	Glu	Lys 2265	ΑΊа	Gln	Аla		Cys 2270	Gln	Thr
Ala		G1n 2275	Gly	Pro	Ala		Ala 2280	His	Cys	Ser	Ser	Arg 2285	٧a٦	Pro	Ile
	Pro 290	Phe	Leu	Leu		Cys 2295	Met	Asn	Ser		Cys 2300	Glu	Phe	Arg	Glu
Leu 2305		Arg	Ala		Cys 2310	Asp	Ser	Leu		Ser 2315	Phe	Glu	Asp	Ala	Cys 2320
Gln	Asn	Gln		Leu 2325	Lys	Pro	Pro		Trp 2330	Arg	Asn	Ser	Ser	Phe 2335	Cys
Pro	Leu		Cys 2340	Pro	Ala	His	Ser	ніs 2345	Tyr	Thr	Asn		Leu 2350	Pro	Ser
Cys		Pro 2355	Ser	Cys	Leu	Asp	Pro 2360	Asp	Ser	Arg		G]u 2365	Gly	ser	Gly
	Lys 2370	٧a٦	Pro	Ala		Cys 2375	Arg	Glu	Gly		11e 2380	Cys	Gln	Pro	Asp
Tyr 2385		Leu	Leu		Asp 2390	Lys	Cys	val	Leu	Arg 2395	Ser	His	Cys	Gly	Cys 2400
Lys	Asp	Ala	Gln	Gly 2405	val	Phe	Ile		А]а 2410	Gly	Lys	Thr	Trp	11e 2415	ser
Glu	Asp		Thr 2420	Gln	Ser	Cys	Thr	Cys 2425	Met	Lys	Glу		Met 2430	Arg	Cys
Trp		Phe 2435	Gln	Cys	Pro		G]y 2440	Thr	Tyr	Cys	Lys	Asn 2445	Ser	Asn	Asp
	ser 2450	Ser	Asn	Cys		Lys 2455	Ile	Ser	Leu		Cys 2460	Pro	Ala	His	Ser
Lys 2465		Thr	Asp		Leu 2470	Pro	Pro	Cys		Pro 2475	Ser	Cys	Ser		Pro 2480
Asp	Gly	His		G]u 2485	Gly	Ile	Ser		Asn 2490	Ala	His	Ser	Asn	Cys 2495	Lys
Glu	Gly		Val 2500	Cys	Gln	Pro	Gly	Tyr 2505	۷a٦	Leu	Arg	Asn	Asp 2510	Lys	Cys
val		Arg 2515	Ile	Glu	Cys	Gly	Cys 2520	Gln				G]y 2525	Gly	Phe	Ile
									Pa	age :	тоэ				

Pro Ala G 2530	ly Lys	Asn Trp	Thr 2535	Ser	Arg	Gly	Cys	Ser 2540	Gln	Ser	Cys	Asp
Cys Met G 2545	lu Gly	Val Ile 2550		Cys	∙Gln		Phe 2555	Gln	Cys	Pro	Ser	G]y 2560
Thr Tyr C		Asp Ile 2565	Glu	Asp	Gly	Thr 2570	Ser	Asn	Cys	Ala	Asn 2575	Ile
Thr Leu G	1n Cys 2580	Pro Ala	His	Ser	Ser 2585	Phe	Thr	Asn	Cys	Leu 2590	Pro	Pro
Cys Gln P 25	ro Ser 95	Cys Ser	Asp	Pro 2600	Glu	Gly	His	Cys	Gly 2605	GТу	Ser	Thr
Thr Lys A 2610	la Pro	Ser Ala	Cys 2615	Gln	Glu	Gly	Cys	Val 2620	Cys	Glu	Pro	Asp
Tyr Val V 2625	al Leu	Asn Asr 2630		Cys	٧a٦	Pro	Arg 2635	Ile	Glu	Cys	Gly	Cys 2640
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Lys Gly C	ys Thr 2660	Gln Th	· Cys	Ala	Cys 2665	٧a٦	Thr	Gly	Thr	11e 2670	His	Cys
Arg Asp P 26	he Gln 75	Cys Pro		Gly 2680	Thr	Tyr	Cys	Lys	Asp 2685	Ile	Lys	Asp
Asp Ala S 2690	er Asn	Cys Th	Glu 2695	Ile	Ile	Leu	Gln	Cys 2700	Pro	Asp	His	Ser
Leu Tyr T 2705	hr His	Cys Let 2710		Ser	Cys		Leu 2715	Ser	Cys	Ser	Asp	Pro 2720
Asp Gly L		Arg Gly 2725	/ Thr	Ser		G1u 2730	Ala	Pro	Ser	Thr	Cys 2735	Lys
Glu Gly C	ys Val 2740	Cys Ası	) Pro	Asp	Tyr 2745	val	Leu	ser	Asn	Asp 2750	Lys	Cys
Val Leu A 27	rg Ile '55	Glu Cys	Gly	Cys 2760	Lys	Asp	Аlа	Gln	G]y 2765	٧a٦	Leu	Ile
Pro Ala G 2770	ily Lys	Thr Tr	11e 2775	Asn	Arg	Gly	Cys	Thr 2780	Gln	Ser	Cys	Ser
Cys Met G 2785	ily Gly	Ala Ile 2790	e Gln )	Cys	Gln	Asn	Phe 2795	Lys	Cys	Pro	Ser	G]u 2800
Ala Tyr C	ys Gln	Asp Met 2805	Glu	Asp	Gly	Asn 2810	ser	Asn	Cys	Thr	Ser 2815	Ile

Pro Leu Gln Cys Pro Ala His Ser His Tyr Thr Asn Cys Leu Pro Thr 2820 2825 2830

Cys Gln Pro Ser Cys Ser Asp Pro Asp Gly His Cys Glu Gly Ser Ser 2835 2840 2845

Thr Lys Ala Pro Ser Ala Cys Lys Glu Gly Cys Val Cys Glu Pro Asp 2850 2855 2860

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Tyr Val Me 2865	et Leu	Asn Asn 2870		Cys	۷a٦	Pro 2	Arg 2875	Ile	Glu	Cys	Gly 2	Cys 2880
Lys Asp Th		Gly Val 2885	Leu	Ile	Pro	Ala 2890	Asp	Lys	Thr	Trp 2	11e 2895	Asn
Arg Gly Cy	/s Thr 2900	Gln Ser	Cys	Thr	Cys 2905	Arg	Gly	Gly	Ala	11e 2910	Gln	Cys
Gln Lys Ty 291		Cys Ser	Ser	G]y 2920	Thr	Tyr	Cys		Asp 2925	Met	Glu	Asp
Asp Ser Se 2930	er Ser		Thr 2935	Ile	Thr	Leu	Gln	Cys 2940	Pro	Ala	His	Ser
His Phe Th 2945	nr Asn	Cys Leu 2950		Pro	Cys		Pro 2955	Ser	Cys	Leu	Asp	ser 2960
Glu Gly H	is Cys	Glu Gly 2965	Ser	Thr	Thr	Lys 2970	Ala	Pro	Ser	Ala	Cys 2975	Gln
Glu Gly Cy	ys Val 2980	Cys Glu	Pro	Asp	Tyr 2985	val	۷al	Leu	Asn	Asn 2990	Lys	Cys
val Pro Ai 299		Glu Cys		Cys 3000	Lys	Asp	Ala		G]y 3005	val	Leu	Ile
Pro Ala As 3010	sp Lys	Thr Trp	lle 3015	Asn	Arg	Gly	Cys	Thr 3020	Gln	Ser	Cys	Thr
Cys Lys G	ly Gly	Ala Ile 3030		Cys	G]n		Phe 3035	Gln	Cys	Pro	Ser	Glu 3040
Thr Tyr Cy		Asp Ile 3045	Glu	Asp	Gly	Asn 3050	Ser	Asn	Cys	Thr	Arg 3055	īle
Ser Leu G	n Cys 3060	Pro Ala	Asn		Asn 3065	Phe	Thr	Ser	Cys	Leu 3070	Pro	Ser
Cys Gln P		Cys Ser		Thr 3080	Asp	val	His	Cys	G]u 3085	Gly	Ser	Ser
Pro Asn Ti 3090	hr Leu	Ser Ser	Cys 3095	Arg	Glu	Gly	Cys	Val 3100	Cys	Gln	Ser	GТу
Tyr Val Lo 3105	eu His	Asn Asp 3110		Cys	Ile	Leu	Arg 3115	Asn	Gln	Cys	Gly	Cys 3120
Lys Asp A		Gly Ala 3125	Leu	Ile	Pro	Glu 3130	Gly	Lys	Thr	Тгр	11e 3135	Thr
Ser Gly C	ys Thr 3140		Cys	Asn	Cys 3145	Thr	Glу	Gly	Аla	11e 3150	Gln	Cys
Gln Asn P		Cys Pro		Lys 3160	Thr	Tyr	Cys		Asp 3165	Leu	Lys	Asp

Gly Ser Ser Asn Cys Thr Asn Ile Pro Leu Gln Cys Pro Ala His Ser 3170 3175 3180

Arg Tyr Thr Asn Cys Leu Pro Ser Cys Pro Pro Leu Cys Leu Asp Pro 3185 3190 3195 3200

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Glu	Gly	Leu		Glu 3205	Gly	Thr	Ser	Pro	Lys 3210	Val	Pro	Ser	Thr	Cys 3215	Arg
Glu	Gly		11e 3220	Cys	Gln	Pro	Gly	Tyr 3225	Leu	Met	His	Lys	Asn 3230	Lys	Cys
val		Arg 3235	Ile	Phe	Cys	Gly	Cys 3240	Lys	Asn	Thr	Gln	Gly 3245	Ala	Phe	Ile
	Ala 3250	Asp	Lys	Thr	Тгр	11e 3255	Ser	Arg	Gly	Cys	Thr 3260	Gln	Ser	Cys	Thr
Cys 326		Ala	Gly		11e 3270	His	Cys	Arg		Phe 3275	Lys	Cys	Pro	Ser	G]y 3280
Thr	Tyr	Cys	Lys	Asn 3285	Gly	Asp	Asn	Gly	Ser 3290	Ser	Asn	Cys	Thr	Glu 3295	IJе
Thr	Leu		Cys 3300	Pro	Thr	Asn	Ser	G]n 3305	Phe	Thr	Asp	Cys	Leu 3310	Pro	Ser
Cys		Pro 3315	Ser	Cys	Ser		Arg 3320	Cys	Glu	٧a٦	Thr	Ser 3325	Pro	ser	٧al
	Ser 3330	Ser	Cys	Arg	Glu	G]y 3335	Cys	Leu	Cys	Asn	ніs 3340	Gly	Phe	٧a٦	Phe
Ser 334	Glu 5	Asp	Lys		Val 3350	Pro	Arg	Thr		Cys 3355	Gly	Cys	Lys	Asp	А1а 3360
Arg	Gly	Ala		11e 3365	Pro	Аlа	Gly		Thr 3370	Тгр	Thr	Ser		Gly 3375	Cys
Thr	Gln		Cys 3380	Аlа	Cys	Val	Glu	G1y 3385	Asn	IJе	G∏n	Cys	G]n 3390	Asn	Phe
Gln		Pro 3395	Pro	Glu	Thr		Cys 3400	Lys	Asp	Asn	Ser	G7u 3405	Gly	ser	Ser
	Cys 3410	Thr	Lys	Ile		Leu 3415	Gln	Cys	Pro		ніs 3420	Thr	Gln	Tyr	Thr
Ser 342	Cys 5	Leu	Pro	Ser	Cys 3430	Leu	Pro	Ser	Cys	Leu 3435	Asp	Pro	Glu	Gly	Leu 3440
Cys	Lys	Asp		Ser 3445	Pro	Lys	val		Ser 3450	Thr	Cys	Lys	Glu	Gly 3455	Cys
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ΑΊа		Cys 3475	Asp	Cys	Lys	Asp	Ala 3480	Gln	Gly	Ala	Leu	11e 3485	Pro	Ala	Gly
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G]y 350		۷al	Gln		G]n 3510	Ser	Ser	Gln		Pro 3515	Pro	Gly	Thr	Tyr	Cys 3520
Lys	Asp	Asn		Asp 3525	Gly	Asn	Ser	Asn	3530	Ala age		Ile	Thr	Leu 3535	Gln

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Ser	Cys	Leu 3555	Asp	Pro	Asp	Gly	Leu 3560	Cys	Lys	Gly	Ala	Ser 3565	Pro	Lys	Val
	Ser 3570	Thr	Cys	Lys	Glu	Gly 3575	Cys	Ile	Cys	Gln	Ser 3580	Gly	Tyr	۷al	Leu
Ser 358!		Asn	Lys	Cys	Leu 3590	Leu	Arg	Asn		Cys 3595	Gly	Cys	Lys		Ala 3600
ніѕ	Gly	Ala		11e 3605	Pro	Glu	Asp		Thr 3610	Trp	٧a٦	Ser		Gly 3615	Cys
Thr	Gln		Cys 3620	٧a٦	Cys	Thr		Gly 3625	Ser	Ile	Gln		Leu 3630	Ser	Ser
Gln		Pro 3635	Pro	Gly	ΑΊа		Cys 3640	Lys	Asp	Asn		Asp 3645	GJy	Ser	Ser
	Cys 3650	Ala	Arg	Ile		Pro 3655	Gln	Cys	Pro		Asn 3660	Ser	His	Tyr	Thr
Asp 366	Cys 5	Phe	Pro	Pro	Cys 3670	Pro	Pro	Ser	Cys	Ser 3675	Asp	Pro	Glu	Gly	ніs 3680
Cys	Glu	Ala		G]y 3685	Pro	Arg	۷al		Ser 3690	Thr	Cys	Arg		Gly 3695	Cys
Leu	Cys		Pro 3700	Gly	Phe	val		Asp 3705	Arg	Asp	Lys		val 3710	Pro	Arg
va1		Cys 3715	Gly	Cys	Lys		А]а 3720	Gln	Gly	Ala		11e 3725	Pro	Ser	GТу
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Ser		Leu 3795	Asp	Pro	Ser		Leu 3800	Cys	Gly	Gly		Ser 3805	Pro	Lys	Glу
	Ser 3810	Thr	Cys	Lys		Gly 3815	Cys	٧al	Cys		Pro 3820	Gly	Tyr	Val	Leu
Asp 382		Asp	Lys	Cys	11e 3830	Leu	Lys	Ile		Cys 3835	Gly	Cys	Arg		Thr 3840
Gln	Gly	Ala		Ile 3845	Pro	Ala	Gly	Lys	Thr 3850	Trp	Leu	Ser		G]y 3855	Cys

Ile Gln Ser Cys Ala Cys Val Glu Gly Thr Ile Gln Cys Gln Asn Phe 3860 3870 Page 169

- Gln Cys Pro Pro Gly Thr Tyr Cys Asn His Asn Asn Asn Cys Ala Lys 3875 3880 3885
- Ile Pro Leu Gln Cys Pro Ala His Ser His Phe Thr Ser Cys Leu Pro 3890 3895 3900
- Ser Cys Pro Pro Ser Cys Ala Asn Leu Asp Gly Ser Cys Glu Gln Thr 3905 3910 3915 3920
- Ser Pro Lys Val Pro Ser Thr Cys Lys Glu Gly Cys Leu Cys Gln Pro 3925 3930 3935
- Gly Tyr Phe Leu Asn Asn Gly Lys Cys Val Leu Gln Thr His Cys Asp 3940 3945 3950
- Cys Lys Asp Ala Glu Gly Gly Leu Val Pro Ala Gly Lys Thr Trp Thr 3955 3960 3965
- Ser Lys Asp Cys Thr Gln Ser Cys Ala Cys Thr Gly Gly Ala Val Gln 3970 3975 3980
- Cys Gln Asn Phe Gln Cys Pro Leu Gly Thr Tyr Cys Lys Asp Ser Gly 3985 3990 3995 4000
- Asp Gly Ser Ser Asn Cys Thr Lys Ile His Lys Gly Ala Met Gly Asp 4005 4010 4015
- Gly Val Leu Met Ala Gly Gly Ile Arg Ala Leu Gln Cys Pro Ala His 4020 4025 4030
- Ser His Phe Thr Ser Cys Leu Pro Ser Cys Pro Pro Ser Cys Ser Asn 4035 4040 4045
- Leu Asp Gly Ser Cys Val Glu Ser Asn Phe Lys Ala Pro Ser Val Cys 4050 4060
- Lys Lys Gly Cys Ile Cys Gln Pro Gly Tyr Leu Leu Asn Asn Asp Lys 4065 4070 4075 4080
- Cys Val Leu Arg Ile Gln Cys Gly Cys Lys Asp Thr Gln Gly Gly Leu 4085 4090 4095
- Ile Pro Ala Gly Arg Thr Trp Ile Ser Ser Asp Cys Thr Lys Ser Cys  $4100 \hspace{1cm} 4105 \hspace{1cm} 4110$
- Ser Cys Met Gly Gly Ile Ile Gln Cys Arg Asp Phe Gln Cys Pro Pro 4115 4120 4125
- Gly Thr Tyr Cys Lys Glu Ser Asn Asp Ser Ser Arg Thr Cys Ala Lys 4130 4135 4140
- Ile Pro Leu Gln Cys Pro Ala His Ser His Tyr Thr Asn Cys Leu Pro 4145 4150 4155 4160
- Ala Cys Ser Arg Ser Cys Thr Asp Leu Asp Gly His Cys Glu Gly Thr 4165 4170 4175
- Ser Pro Lys Val Pro Ser Pro Cys Lys Glu Gly Cys Leu Cys Gln Pro 4180 4185 4190
- Gly Tyr Val Val His Asn His Lys Cys Val Leu Gln Ile His Cys Gly 4195 4200 4205 Page 170

Cys Lys Asp Ala Gln Gly Gly Phe Val Pro Ala Gly Lys Thr Trp Ile 4210 4215 4220 Ser Arg Gly Cys Thr Gln Ser Cys Ala Cys Val Gly Gly Ala Val Gln 4225 4230 4235 4240 Cys His Asn Phe Thr Cys Pro Thr Gly Thr Gln Cys Gln Asn Ser Ser 4245 4250 4255 Cys Ser Lys Ile Thr Val Gln Cys Pro Ala His Ser Gln Tyr Thr Thr 4260 4265 4270 Cys Leu Pro Ser Cys Leu Pro Ser Cys Phe Asp Pro Glu Gly Leu Cys 4280 Gly Gly Ala Ser Pro Arg Ala Pro Ser Thr Cys Arg Glu Gly Cys Val 4290 4295 4300 Cys Glu Ala Asp Tyr Val Leu Arg Glu Asp Lys Cys Val Leu Arg Thr 4305 4310 4315 4320 Gln Cys Gly Cys Lys Asp Ala Gln Gly Asp Leu Ile Pro Ala Asn Lys 4325 4330 4335 Thr Trp Leu Thr Arg Gly Cys Ala Gln Lys Cys Thr Cys Lys Gly Gly 4340 4345 4350 Asn Ile His Cys Trp Asn Phe Lys Cys Pro Leu Gly Thr Glu Cys Lys 4355 4360 4365 Asp Ser Val Asp Gly Gly Ser Asn Cys Thr Lys Ile Ala Leu Gln Cys 4370 4375 4380 Pro Ala His Ser His His Thr Tyr Cys Leu Pro Ser Cys Ile Pro Ser Cys Ser Asn Val Asn Asp Arg Cys Glu Ser Thr Ser Leu Gln Arg Pro 4405 4410 4415 Ser Thr Cys Ile Glu Gly Cys Leu Cys His Ser Gly Phe Val Phe Ser 4420 4425 4430 Lys Asp Lys Cys Val Pro Arg Thr Gln Cys Gly Cys Lys Asp Ser Gln
4435 4440 4445 Gly Thr Leu Ile Pro Ala Gly Lys Asn Trp Ile Thr Thr Gly Cys Ser 4450 4455 4460 Gln Arg Cys Thr Cys Thr Gly Gly Leu Val Gln Cys His Asp Phe Gln 4465 Cys Pro Ser Gly Ala Glu Cys Gln Asp Ile Glu Asp Gly Asn Ser Asn 4485 4490 4495 Cys Val Glu Ile Thr Val Gln Cys Pro Ala His Ser His Tyr Ser Lys 4500 4505 4510 Cys Leu Pro Pro Cys Gln Pro Ser Cys Ser Asp Pro Asp Gly His Cys 4515 4520 4525

Glu Gly Thr Ser Pro Glu Ala Pro Ser Thr Cys Glu Glu Gly Cys Val

4540 Page 171

4535

Cys Glu Pro Asp Tyr Val Leu Ser Asn Asp Lys Cys Val Pro Ser Ser 4545 4550 4555 4560 Glu Cys Gly Cys Lys Asp Ala His Gly Val Leu Ile Pro Glu Ser Lys 4565 4570 4575 Thr Trp Val Ser Arg Gly Cys Thr Lys Asn Cys Thr Cys Lys Gly Gly 4580 4585 4590 Thr Val Gln Cys His Asp Phe Ser Cys Pro Thr Gly Ser Arg Cys Leu 4595 4600 4605 Asp Asn Asn Glu Gly Asn Ser Asn Cys Val Thr Tyr Ala Leu Lys Cys 4610 4615 4620 Pro Ala His Ser Leu Tyr Thr Asn Cys Leu Pro Ser Cys Leu Pro Ser Cys Ser Asp Pro Glu Gly Leu Cys Gly Gly Thr Ser Pro Glu Val Pro 4645 4650 4655 Ser Thr Cys Lys Glu Gly Cys Ile Cys Gln Ser Gly Tyr Val Leu His 4660 4665 4670 Lys Asn Lys Cys Met Leu Arg Ile His Cys Asp Cys Lys Asp Phe Gln 4675 4680 4685 Gly Ser Leu Ile Lys Thr Gly Gln Thr Trp Ile Ser Ser Gly Cys Ser 4690 4695 4700 Lys Ile Cys Thr Cys Lys Gly Gly Phe Phe Gln Cys Gln Ser Tyr Lys 4705 4710 4715 4720 Cys Pro Ser Gly Thr Gln Cys Glu Glu Ser Glu Asp Gly Ser Ser Asn 4725 4730 4735 Cys Val Ser Ser Thr Met Lys Cys Pro Ala Asn Ser Leu Tyr Thr His 4740 4745 4750 Cys Leu Pro Thr Cys Leu Pro Ser Cys Ser Asn Pro Asp Gly Arg Cys 4755 4760 4765 Glu Gly Thr Ser His Lys Ala Pro Ser Thr Cys Arg Glu Gly Cys Val 4770 4775 4780 Cys Gln Pro Gly Tyr Leu Leu Asn Lys Asp Thr Cys Val His Lys Asn 4785 4790 4705 Gln Cys Gly Cys Lys Asp Ile Arg Gly Asn Ile Ile Pro Ala Gly Asn 4805 4810 4815 Thr Trp Ile Ser Ser Asp Cys Thr Gln Ser Cys Ala Cys Thr Asp Gly Val Ile Gln Cys Gln Asn Phe Val Cys Pro Ser Gly Ser His Cys Gln Cys Thr Ile Phe Gly Asp Pro Tyr Tyr Leu Thr Phe Asp Gly Phe Thr 4870 4875 Page 172

- Tyr His Phe Leu Gly Arg Met Asn Tyr Tyr Leu Ile Lys Thr Val Asp 4885 4890 4895
- Lys Leu Pro Arg Gly Ile Glu Pro Leu Ile Met Glu Gly Arg Asn Lys 4900 4905 4910
- Ile Ser Pro Lys Gly Ser Ser Thr Leu His Glu Val Thr Thr Ile Val 4915 4920 4925
- Tyr Gly Tyr Lys Ile Gln Leu Gln Glu Glu Leu Val Val Leu Val Asn 4930 4935 4940
- Asp Glu Lys Val Ala Val Pro Tyr Asn Pro Asn Glu His Leu Arg Val 4945 4950 4955 4960
- Met Leu Arg Ala Gln Arg Leu Leu Leu Val Thr Asp Phe Glu Met Val 4965 4970 4975
- Leu Asp Phe Asp Gly Lys His Ser Ala Val Ile Ser Leu Pro Thr Thr 4980 4985 4990
- Tyr Arg Gly Leu Thr Arg Gly Leu Cys Gly Asn Tyr Asp Arg Asp Gln 4995 5000 5005
- Ser Asn Glu Leu Met Leu Pro Ser Gly Val Leu Thr Ser Asn Val His 5010 5015 5020
- Val Phe Gly Asn Ser Trp Glu Val Lys Ala Gln His Ala Phe Phe Arg 5025 5030 5035 5040
- Phe Pro Arg Ala Leu Pro Glu Asp Glu Glu Arg Asp Glu Glu Pro Asp 5045 5050 5055
- Leu Leu Gln Ser Glu Cys Ser Gln Glu Gln Thr Ala Leu Ile Ser Ser 5060 5065 5070
- Thr Gln Ala Cys Arg Val Leu Val Asp Pro Gln Gly Pro Phe Ala Ala 5075 5080 5085
- Cys His Gln Ile Ile Ala Pro Glu Pro Phe Glu Gln Arg Cys Met Leu 5090 5095 5100
- Asp Met Cys Thr Gly Trp Lys Thr Lys Glu Glu Glu Glu Leu Arg Cys 5105 5110 5120
- Arg Val Leu Ser Gly Tyr Ala Ile Ile Cys Gln Glu Ala Gly Ala Asn 5125 5130 5135
- Met Thr Gly Trp Arg Asp His Thr His Cys Ala Met Thr Cys Pro Ala 5140 5145 5150
- Asn Thr Val Tyr Gln Arg Cys Met Thr Pro Cys Pro Ala Ser Cys Ala 5155 5160 5165
- Lys Phe Val Thr Pro Lys Val Cys Glu Gly Pro Cys Val Glu Gly Cys 5170 5180
- Ala Ser Leu Pro Gly Tyr Ile Tyr Ser Asp Thr Gln Ser Leu Pro Val 5185 5190 5195 . 5200
- Thr His Cys Gly Cys Thr Ala Asp Gly Ile Tyr Tyr Lys Leu Gly Asp 5205 5210 5215 Page 173

Ser Phe Val Thr Asn Asp Cys Ser Gln His Cys Thr Cys Ala Ser Gln Gly Ile Leu Leu Cys Glu Pro Tyr Gly Cys Arg Ala Gly Glu Ser Cys 5235 Leu Cys Glu Pro Tyr Gly Cys Arg Ala Gly Glu Ser Cys Met Val Ala Asn Phe Thr Arg Gly Cys Phe Gln Asp Ser Pro Cys Leu 5250 Asn Pro Cys His Asn Asp Gly Arg Cys Glu Glu Gln Gly Ala Thr 5265 Phe Ile Cys His Cys Asp Phe Gly Tyr Gly Gly Glu Phe Cys Thr Glu Pro Gln Asp Ile Thr Thr Arg Lys Lys Ile Glu Ala Ser Ser Leu Val Ala Ile Leu Pro Gly Val Leu Val Met Val Leu Val Pro Val Leu Leu Pro Arg Val Tyr Met Ala Thr Arg Thr Thr Met Gly Arg Arg 5330 Arg Met Lys Arg Lys Glu Lys Lys Leu Leu Arg Gln Ser Arg Leu Arg 5345 Leu Glu Asp Ala Asp Val Pro Glu Pro Thr Phe Lys Ala Thr Glu Phe 5375

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<212> PRT
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Pro Met Val His Thr Ser Arg Glu Asp Ser Ile Leu Ser Lys Cys Asp
Phe Glu Asp Asn Ser Arg Pro Phe Cys Asp Trp Ser Gln Met Ser Ala
50
Asp Asp Gly Asp Trp Ile Arg Thr Thr Gly Pro Ser Leu Thr Gly Thr
65
Ser Gly Pro Pro Gly Gly Tyr Pro Asn Gly Glu Gly Tyr Tyr Leu His
Met Asp Pro Lys Thr Phe Pro Gln Gly Gly Val Ala Arg Leu Arg Ser
Pro Asp Ile Trp Glu Gln Gly Pro Leu Cys Val His Phe Ala Phe His
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<sup>&</sup>lt;213> Mus musculus

Met Phe Gly Leu Ser Trp Gly Ala Gln Leu Arg Leu Leu Leu Leu Arg 130 135 140 Gly Arg Lys His Leu Arg Pro Tyr Val Leu Trp Lys His Val Asn Thr 145 150 155 160 Gln Ser Pro Ser Trp Met Pro Thr Thr Val Thr Val Pro Ala Asp His 165 170 175 Asp Ile Pro Ser Trp Leu Met Phe Glu Gly Met Arg Gly Asn Thr Ala 180 185 190 Tyr Leu Asp Ile Ser Leu Asp Gly Leu Ser Ile Gln Arg Gly Thr Cys
200 205 Asn Gln Val Cys Met Ser Gln Met Cys Thr Phe Asp Thr Leu Asn Asp 210 220 Leu Cys Gly Trp Ser Trp Val Pro Thr Ala Thr Gly Ala Lys Trp Thr 225 230 235 240 Gln Lys Lys Gly Pro Thr Gly Lys Gln Gly Val Gly Pro Ala Glu Asp 245 250 255 Phe Ser Asn Pro Gly Asn Gly Tyr Tyr Met Leu Leu Asp Ser Thr Asn 260 265 270 Ala Arg Pro Gly Gln Lys Ala Val Leu Leu Ser Pro Leu Ser His Ser 275 280 285 Arg Gly Cys Met Thr Leu Ser Phe His Tyr Ile Met His Gly Gln Gly 290 295 300 His Glu Glu Gly Leu Phe Val Tyr Ala Thr Phe Leu Gly Asn Ile Arg 305 310 315 320 Lys Tyr Thr Leu Phe Ser Gly His Pro Gly Pro Asp Trp Gln Ala Val 325 330 335 Ser Val Asn Tyr Thr Gly Gln Gly Gln Ile Gln Phe Met Val Val Gly 340 345 350Met Phe Gly Asn Ile Pro Glu Pro Ala Ile Ala Val Asp Ala Ile Ser 355 360 365 Ile Ala Pro Cys Gly Glu Ser Phe Pro Gln Cys Asp Phe Glu Asp Arg 370 375 380 Val His Pro Phe Cys Asp Trp Asn Gln Val Tyr Gly Asp Met Gly His 385 390 400 Trp Ser Trp Gly Ser Lys Ser Val Pro Thr Leu Ile Ala Gly Ser Pro 405 410 415 Arg Glu Phe Pro Tyr Gly Gly Glu His Tyr Ile Phe Phe Asp Ser Val 420 425 430 Lys Leu Ser Gln Glu Gly Gln Ser Ala Arg Leu Val Ser Pro Pro Phe 435 440 445 Cys Ala Pro Gly Asp Ile Cys Val Glu Phe Ala Tyr His Met Tyr Gly Page 175

Leu Gly Lys Gly Thr Thr Leu Lys Leu Leu Gly Ser Pro Ala Gly 465 470 480 470 Ser Phe Pro Ile Pro Leu Trp Asn Arg Val Gly Ser Gln Ser Ser Gly 485 490 495 Trp Met Asn Ser Ser Val Thr Ile Pro Lys Gly Tyr Gln Gln Pro Met  $500 \hspace{1.5cm} 505 \hspace{1.5cm} 510$ Gln Leu Phe Ile Glu Ala Thr Arg Gly Thr Ser Thr Ala Phe Val Val 515 520 525 Ala Leu Asn Phe Ile Leu Ile Ser His Gly Pro Cys Arg Val Leu Leu 530 540 Gln Thr Glu Ile Pro Ser Ser Pro Leu Leu Pro Pro Thr Gly Pro Ser 545 550 560 Glu Ser Thr Val Pro Thr Leu Pro Met Glu Gln Pro Thr Ser Pro Thr 565 570 575 Lys Ala Thr Thr Val Thr Ile Glu Ile Pro Thr Thr Pro Thr Glu Glu 580 590 Ala Thr Ile Pro Thr Glu Thr Thr Thr Val Pro Thr Glu Val Ile Asn 595 600 605 Val Ser Pro Lys Glu Thr Ser Ile Pro Pro Glu Val Thr Ile Pro Thr Glu Val Ile Thr Val Ser Pro Glu Glu Ile Ile Ser Pro Thr Glu Val 625 630 635 640 Thr Pro Val Pro Thr Asp Val Thr Ala Ala Tyr Val Glu Ala Thr Asn 645 650 655 Ala Ser Pro Glu Glu Thr Ser Val Pro Pro Glu Val Thr Ile Leu Thr Glu Val Thr Thr Val Ser Pro Glu Glu Thr Thr Val Pro Thr Glu Val 675 680 685 Pro Ile Val Leu Ile Glu Ala Thr Ala Phe Pro Thr Gly Glu Thr Thr 690 695 700 Leu Tyr Thr Glu Val Pro Thr Val Pro Thr Glu Val Thr Gly Val His 705 710 715 720 Thr Glu Val Thr Asn Val Ser Pro Glu Glu Thr Ser Val Pro Thr Glu 725 730 735 Glu Thr Ile Ser Thr Glu Val Thr Thr Val Ser Pro Glu Glu Thr Thr 740 745 750 Leu Pro Thr Glu Val Pro Thr Val Ser Thr Glu Val Thr Asn Val Ser 755 760 765 Pro Glu Glu Thr Ser Val Pro Pro Glu Glu Thr Ile Leu Thr Glu Ile 770 775 780 Thr Thr Val Ser Pro Glu Glu Thr Val Phe Pro Thr Glu Gly Thr Thr Page 176

Leu Pro Thr Glu Val Leu Thr Val Pro Ile Glu Val Thr Thr Phe Pro 810 815 805 Thr Gly Glu Thr Thr Val Pro Thr Glu Val Pro Thr Val Ser Thr Glu Met Thr Gly Val His Thr Glu Val Thr Thr Val Phe Pro Glu Glu Thr 835 840 845 Ser Ile Pro Thr Glu Val Ala Thr Val Leu Pro Ala Ser Ile Pro Pro Glu Glu Thr Thr Thr Pro Thr Glu Val Thr Thr Thr Pro Pro Glu Glu 875 Thr Thr Ile Pro Ala Glu Val Thr Thr Val Pro Pro Val Ser Ile Pro 890 Ser Glu Glu Thr Thr Pro Thr Glu Val Thr Thr Pro Pro Glu Glu Thr Thr Ile Pro Ala Glu Val Thr Thr Val Pro Pro Val Ser Ile 920 Pro Ser Glu Glu Thr Thr Thr Pro Thr Glu Val Thr Thr Thr Pro Pro 935 Glu Glu Thr Thr Ile Pro Ala Glu Val Thr Thr Val Pro Pro Val Ser 960 950 Ile Pro Ser Glu Glu Thr Thr Ile Pro Thr Glu Val Thr Thr Val Pro 970 Pro Glu Glu Thr Thr Ile Pro Ala Glu Val Thr Thr Val Pro Pro Val 985 Ser Ile Pro Ser Glu Glu Thr Thr Ile Pro Thr Glu Val Thr Thr Val 1000 Pro Pro Glu Glu Thr Thr Ile Pro Ala Glu Val Thr Thr Pro Pro 1015 1020 Glu Glu Thr Thr Ile Pro Thr Glu Val Thr Thr Val Pro Pro Ala Ser 1030 1035 Ile Pro Pro Glu Glu Thr Ala Ser Leu Thr Glu Val Thr Thr Thr Pro 1050 1045 Pro Glu Glu Thr Thr Thr Pro Thr Glu Val Thr Thr Val Pro Pro Glu 1065 1070 Lys Thr Thr Ile Pro Thr Glu Val Thr Thr Val Pro Pro Ala Ser Ile 1085 1080 Phe Pro Glu Glu Thr Thr Val Pro Pro Glu Glu Thr Thr Ile Ala Ser 1095 1100 Glu Glu Thr Thr Val Ser Thr Gln Glu Thr Thr Leu Leu Thr Glu Gln 1115 1105 1110 Ser Ala Val Thr Gln Thr Ser Ile Ala Cys Arg Pro Pro Cys Pro Ser

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Pro Pro Leu Met Pro Ile Gly Pro Leu Leu Ser Lys Pro Pro Gly Val 1140 1145 1150

Ser Met Phe Ser Leu Ala Pro Thr Thr Gly Val Ser Thr Thr Glu Ser 1155 1160 1165

Cys Pro Pro Asn Ala His Ile Glu Leu Cys Ala Cys Pro Ala Ser Cys 1170 1175 1180

Glu Ser Pro Lys Pro Ser Cys Gln Pro Pro Cys Ile Pro Gly Cys Val 1185 1190 1195 1200

Cys Asn Pro Gly Phe Leu Phe Ser Asn Asn Gln Cys Ile Asn Glu Ser 1205 1210 1215

Ser Cys Asn Cys Pro Tyr Asn Asn Lys Tyr Tyr Lys Pro Gly Glu Glu 1220 1225 1230

Trp Phe Thr Pro Asn Cys Thr Glu Arg Cys Arg Cys Leu Pro Gly Ser 1235 1240 1245

Leu Met Glu Cys Gln Ile Ser Gln Cys Gly Thr His Thr Val Cys Gln 1250 1255 1260

Leu Lys Ser Asp Gln Tyr Gln Cys Glu Pro Tyr Gly Lys Ala Thr Cys 1265 1270 1275 1280

Leu Val Tyr Gly Asp Leu His Phe Val Thr Phe Asp Glu Arg His Ile 1285 1290 1295

Gly Phe Thr Gly Thr Cys Thr Tyr Ile Leu Thr Gln Thr Cys Ser Asn  $1300 \hspace{1cm} 1305 \hspace{1cm} 1310$ 

Ser Thr Asp His Phe Phe Arg Ile Thr Ala Asn Thr Glu Glu Arg Gly 1315 1320 1325

Val Glu Gly Val Ser Cys Leu Asp Lys Val Val Ile Ser Leu Pro Glu 1330 1340

Thr Thr Val Thr Met Ile Ser Gly Arg His Thr Leu Ile Gly Asp Gln 1345 1350 1360

Glu Val Thr Leu Pro Ala Ile Leu Ser Asp Asp Thr Tyr Val Gly Leu 1365 1370 1375

Ser Gly Arg Phe Val Glu Leu Arg Thr Thr Phe Gly Leu Arg Val Arg 1380 1385 1390

Trp Asp Gly Asp Gln Gln Leu Phe Val Thr Val Ser Ser Thr Phe Ser 1395 1400 1405

Gly Lys Leu Cys Gly Phe Cys Gly Asn Tyr Asp Gly Asp Ser Ser Asn 1410 1415 1420

Asp Asn Leu Lys Ser Asp Gly Met Met Thr His Asp Glu Glu Glu Leu 1425 1430 1435 1440

Arg Leu Ser Trp Gln Val Glu Glu Asp Glu Asp Lys Asp Trp Val Ser 1445 1450 1455

Ser Arg Cys Gln Lys Lys Lys Asn Pro Pro Ser Cys Asp Ala Ala Leu Page 178

1470

Gly Ser Thr Met Ser Gly Pro Lys Leu Cys Gly Gln Leu Val Asn Pro 1480

Ser Gly Pro Phe Glu Ala Cys Leu Leu His Leu Lys Ala Ser Ser Phe 1490 1495 1500

Leu Asp Asn Cys Val Thr Asp Met Cys Ser Phe Gln Gly Leu Gln Gln

Lys Leu Cys Ala His Met Ser Ala Met Thr Ala Thr Cys Gln Asp Ala 1530

Gly Tyr Pro Val Lys Pro Trp Arg Glu Pro Gln Phe Cys Pro Leu Val 1540 1545 1550

Cys Pro Lys Asn Ser Arg Tyr Ser Leu Cys Ala Lys Pro Cys Pro Glu 1555 1560 1565

Thr Cys His Pro Ile Ser Thr Thr Gln His Cys Ser Asp Lys Cys Val

Glu Gly Cys Glu Cys Asp Pro Gly Phe Ile Leu Ser Gly Ser Glu Cys 1585 1590 1595 1600

Val Pro Ser Ser Gln Cys Gly Cys Thr Ser Phe Gln Gly Arg Tyr Phe  $1605 \hspace{1cm} 1610 \hspace{1cm} 1615$ 

Lys Val Gln Glu Gln Trp Phe Asn Pro Asp Cys Lys Glu Ile Cys Thr 1625

Cys Glu Ser His Asn His Ile Leu Cys Lys Pro Trp Lys Cys Lys Ala 1635 1640 1645

Gln Glu Ala Cys Ser Tyr Lys Asn Gly Val Leu Gly Cys His Ala Gln 1650 1655 1660

Gly Ala Ala Thr Cys Met Val Ser Gly Asp Pro His Tyr Leu Thr Phe

Asp Gly Ala Leu His His Phe Met Gly Thr Cys Thr Tyr Val Leu Thr

Gln Pro Cys Trp Ser Lys Ser Gln Glu Asn Asn Phe Val Val Ser Ala 1700 1705 1710

Thr Asn Glu Ile His Asp Gly Asn Leu Glu Val Ser Tyr Val Lys Ala 1715 1720 1725

Val His Val Gln Val Phe Asp Leu Lys Ile Ser Met Phe Lys Gly Gln 1730 1735 1740

Lys Val Val Leu Asn Asn Gln Arg Val Val Leu Pro Val Trp Pro Ser 1745 1750 1755 1760

Gln Gly Arg Val Thr Ile Arg Leu Ser Gly Ile Phe Val Leu Leu Tyr 1765 1770 1775

Thr Asn Phe Gly Leu Gln Val Arg Tyr Asp Gly Arg His Leu Val Glu 1780 1785 1790

val Thr val Pro Ser Ser Tyr Thr Gly Ser Leu Cys Gly Leu Cys Gly

Asn Tyr Asn Asn Asn Ser Met Asp Asp Asn Leu Arg Val Asp Met Lys 1810 1815 1820

Pro Ala Gly Asn Ser Leu Leu Leu Gly Ala Ala Trp Lys Ile Leu Glu 1825 1830 1835 1840

Ala Ser Asp Pro Gly Cys Phe Leu Val Gly Gly Lys Pro Ser Arg Cys 1845 1850 1855

Ala Asp Ser Asp Met Asp Asp Val Trp Thr Lys Lys Cys Ala Ile Leu 1860 1865 1870

Met Asn Pro Leu Gly Pro Phe Ser Asn Cys His Glu Ala Val Pro Pro 1875 1880 1885

Gln Ala Ser Phe Ser Ser Cys Val Tyr Gly Gln Cys Glu Thr Asn Gly 1890 1895 1900

Asp Asn Leu Thr Leu Cys His Ser Leu Gln Ala Tyr Ala Ser Leu Cys 1905 1910 1915 1920

Ala Gln Ala Gly Gln Val Thr Trp Arg Asn Ser Thr Phe Cys Pro 1925 1930 1935

Met Arg Cys Pro Pro Arg Ser Ser Tyr Asn Pro Cys Ala Asn Ser Cys 1940 1945 1950

Pro Ala Thr Cys Leu Thr Leu Ser Thr Pro Arg Asp Cys Pro Thr Leu 1955 1960 1965

Pro Cys Val Glu Gly Cys Glu Cys Gln Ser Gly His Ile Leu Ser Gly 1970 1975 1980

Thr Thr Cys Val Pro Leu Arg Gln Cys Gly Cys Ser Asp Gln Asp Gly 1985 1990 1995 2000

Ser Tyr His Leu Leu Gly Glu Ser Trp Tyr Thr Glu Lys Thr Cys Thr 2005 2010 2015

Thr Leu Cys Thr Cys Ser Ala His Ser Asn Ile Thr Cys Ser Pro Thr 2020 2025 2030

Ala Cys Lys Ala Asn His Val Cys Leu Arg Gln Glu Gly Leu Leu Arg 2035 2040 2045

Cys Ala Ala Glu Met Gly Glu Cys Arg Ile Ser Glu Asp Ser Gln Ile 2050 2055 2060

Val Ser Phe Asp Asp His Ser His Pro Ile Gln Asp Thr Cys Thr Tyr 2065 2070 2075 2080

Ile Leu Val Lys Val Cys His Pro Asn Thr Asn Met Pro Phe Phe Met 2085 2090 2095

Ile Ser Ala Lys Thr Asp Ile Asn Thr Asn Gly Lys Asn Lys Thr Phe  $2100 \hspace{1cm} 2105 \hspace{1cm} 2110$ 

Gly Val Tyr Gln Leu Tyr Ile Asp Ile Phe Asn Phe His Ile Thr Leu 2115 2120 2125

Gln Lys Asp His Leu Val Leu Ile Ser Leu Ile Asn Asp Ser Ile Val Page 180 Thr Leu Pro Thr Thr His Ile Pro Gly Val Ser Val Met Thr Glu 2145 2150 2155 2160

Asp Val Tyr Thr Ile Val Thr Ile Lys Asp Glu Ile Gln Val Lys Phe 2165 2170 2175

Glu Ser Asn Asn Phe Leu Asp Val Lys Ile Pro Ala Ser Ser Asn Gly 2180 2185 . 2190

Lys Val Cys Gly Val Cys Gly Asn Phe Asn Gly Glu Glu Asp Glu 2195 2200 2205

Leu Met Thr Pro Ser Gly Glu Leu Ala Glu Asp Glu Gln Glu Phe Met 2210 2215 2220

Gly Gln Asn Leu Gln Val Glu Gln Gln Glu Ile Met Asn Gly Lys Cys 2245 2250 2255

Arg Pro Ile Asp Phe Glu Lys Ala Gln Ala Asn Cys Gln Thr Ala Leu 2260 2265 2270

Gln Gly Pro Ala Trp Ala His Cys Ser Ser Arg Val Pro Ile Lys Pro 2275 2280 2285

Phe Leu Leu Lys Cys Met Asn Ser Phe Cys Glu Phe Arg Glu Leu Phe 2290 2300

Arg Ala Leu Cys Asp Ser Leu Gln Ser Phe Glu Asp Ala Cys Gln Asn 2305 2310 2315 2320

Gln Gly Leu Lys Pro Pro Ile Trp Arg Asn Ser Ser Phe Cys Pro Leu 2325 2330 2335

Glu Cys Pro Ala His Ser His Tyr Thr Asn Cys Leu Pro Ser Cys Pro 2340 2345 2350

Pro Ser Cys Leu Asp Pro Asp Ser Arg Cys Glu Gly Ser Gly His Lys 2355 2360 2365

Val Pro Ala Thr Cys Arg Glu Gly Cys Ile Cys Gln Pro Asp Tyr Val 2370 2375 2380

Leu Leu Asn Asp Lys Cys Val Leu Arg Ser His Cys Gly Cys Lys Asp 2385 2390 2395 2400

Ala Gln Gly Val Phe Ile Pro Ala Gly Lys Thr Trp Ile Ser Glu Asp 2405 2410 2415

Cys Thr Gln Ser Cys Thr Cys Met Lys Gly Ser Met Arg Cys Trp Asp 2420 2425 2430

Phe Gln Cys Pro Pro Gly Thr Tyr Cys Lys Asn Ser Asn Asp Gly Ser 2435 2440 2445

Ser Asn Cys Val Lys Ile Ser Leu Gln Cys Pro Ala His Ser Lys Phe 2450 2455 2460

Thr Asp Cys Leu Pro Pro Cys His Pro Ser Cys Ser Asp Pro Asp Gly Page 181 His Cys Glu Gly Ile Ser Thr Asn Ala His Ser Asn Cys Lys Glu Gly 2485 2490 2495

Cys Val Cys Gln Pro Gly Tyr Val Leu Arg Asn Asp Lys Cys Val Leu 2500 2505 2510

Arg Ile Glu Cys Gly Cys Gln His Thr Gln Gly Gly Phe Ile Pro Ala 2515 2520 2525

Gly Lys Ser Trp Thr Ser Arg Gly Cys Ser Gln Ser Cys Asp Cys Met 2530 2540

Glu Gly Val Ile Arg Cys Gln Asn Phe Gln Cys Pro Ser Gly Thr Tyr 2545 2550 2560

Cys Gln Asp Ile Glu Asp Gly Thr Ser Asn Cys Ala Asn Ile Thr Leu 2565 2570 2575

Gln Cys Pro Ala His Ser Ser Phe Thr Asn Cys Leu Pro Pro Cys Gln 2580 2585 2590

Pro Ser Cys Ser Asp Pro Glu Gly His Cys Gly Gly Ser Thr Thr Lys 2595 2600 2605

Ala Pro Ser Ala Cys Gln Glu Gly Cys Val Cys Glu Pro Asp Tyr Val 2610 2615 2620

Val Leu Asn Asn Lys Cys Val Pro Arg Ile Glu Cys Gly Cys Lys Asp 2625 2630 2635 2640

Ala Gln Gly Val Leu Ile Pro Ala Asp Lys Ile Trp Ile Asn Lys Gly 2645 2650 2655

Cys Thr Gln Thr Cys Ala Cys Val Thr Gly Thr Ile His Cys Arg Asp 2660 2665 2670

Phe Gln Cys Pro Ser Gly Thr Tyr Cys Lys Asp Ile Lys Asp Asp Ala 2675 2680 2685

Ser Asn Cys Thr Glu Ile Thr Leu Gln Cys Pro Asp His Ser Leu Tyr 2690 2695 2700

Thr His Cys Leu Pro Ser Cys Leu Pro Ser Cys Ser Asp Pro Asp Gly 2705 2710 2715 2720

Leu Cys Arg Gly Thr Ser Pro Glu Ala Pro Ser Thr Cys Lys Glu Gly 2725 2730 2735

Cys Val Cys Asp Pro Asp Tyr Val Leu Ser Asn Asp Lys Cys Val Leu 2740 2745 2750

Arg Ile Glu Cys Gly Cys Lys Asp Ala Gln Gly Val Leu Ile Pro Ala 2755 2760 2765

Gly Lys Thr Trp Ile Asn Arg Gly Cys Thr Gln Ser Cys Ser Cys Met 2770 2780

Gly Gly Ala Ile Gln Cys Gln Asn Phe Lys Cys Pro Ser Glu Ala Tyr 2785 2790 2795 2800

Cys Gln Asp Leu Glu Asp Gly Asn Ser Asn Cys Thr Ser Ile Pro Leu Page 182

Gln Cys Pro Ala His Ser His Tyr Thr Asn Cys Leu Pro Thr Cys Gln 2820 2825 2830

Pro Ser Cys Ser Asp Pro Asp Gly His Cys Glu Gly Ser Ser Thr Lys 2835 2840 2845

Ala Pro Ser Ala Cys Lys Glu Gly Cys Val Cys Glu Pro Asp Tyr Val 2850 2855 2860

Met Leu Asn Asn Lys Cys Val Pro Arg Ile Glu Cys Gly Cys Lys Asp 2865 2870 2875 2880

Thr Gln Gly Val Leu Ile Pro Ala Asp Lys Thr Trp Ile Asn Arg Gly 2885 2890 2895

Cys Thr Gln Ser Cys Thr Cys Lys Gly Gly Ala Ile Gln Cys Gln Lys 2900 2905 2910

Tyr His Cys Ser Ser Gly Thr Tyr Cys Lys Asp Met Glu Asp Asp Ser 2915 2920 2925

Ser Ser Cys Ala Thr Ile Thr Leu Gln Cys Pro Ala His Ser His Phe 2930 2935 2940

Thr Asn Cys Leu Pro Pro Cys Gln Pro Ser Cys Leu Asp Ser Glu Gly 2945 2950 2955 2960

His Cys Glu Gly Ser Thr Thr Lys Ala Pro Ser Ala Cys Gln Glu Gly 2965 2970 2975

Cys Val Cys Glu Pro Asp Tyr Val Val Leu Asn Asn Lys Cys Val Pro 2980 2985 2990

Arg Ile Glu Cys Gly Cys Lys Asp Ala Gln Gly Val Leu Ile Pro Ala 2995 3000 3005

Asp Lys Thr Trp Ile Asn Arg Gly Cys Thr Gln Ser Cys Thr Cys Lys 3010 3015 3020

Gly Gly Ala Ile Gln Cys Gln Lys Phe Gln Cys Pro Ser Glu Thr Tyr 3025 3030 3035 3040

Cys Lys Asp Ile Glu Asp Gly Asn Ser Asn Cys Thr Arg Ile Ser Leu 3045 3050 3055

Gln Cys Pro Ala Asn Ser Asn Phe Thr Ser Cys Leu Pro Ser Cys Gln 3060 3065 3070

Pro Ser Cys Ser Asn Thr Asp Val His Cys Glu Gly Ser Ser Pro Asn 3075 3080 3085

Thr Leu Ser Ser Cys Arg Glu Gly Cys Val Cys Gln Ser Gly Tyr Val 3090 3095 3100

Leu His Asn Asp Lys Cys Ile Leu Arg Asn Gln Cys Gly Cys Lys Asp 3105 3110 3115 3120

Ala Gln Gly Ala Leu Ile Pro Glu Gly Lys Thr Trp Ile Thr Ser Gly 3125 3130 3135

Cys Thr Gln Ser Cys Asn Cys Thr Gly Gly Ala Ile Gln Cys Gln Asn Page 183

Phe Gln Cys Pro Leu Lys Thr Tyr Cys Lys Asp Leu Lys Asp Gly Ser 3155 3160 3165

Ser Asn Cys Thr Asn Ile Pro Leu Gln Cys Pro Ala His Ser Arg Tyr 3170 3175 3180

Thr Asn Cys Leu Pro Ser Cys Pro Pro Ser Cys Leu Asp Pro Glu Gly 3185 3190 3195 3200

Leu Cys Glu Gly Thr Ser Pro Lys Val Pro Ser Thr Cys Arg Glu Gly 3205 3210 3215

Cys Ile Cys Gln Pro Gly Tyr Leu Met His Lys Asn Lys Cys Val Leu 3220 3225 3230

Arg Ile Phe Cys Gly Cys Lys Asn Thr Gln Gly Ala Phe Ile Ser Ala 3235 3240 3245

Asp Lys Thr Trp Ile Ser Arg Gly Cys Thr Gln Ser Cys Thr Cys Ser 3250 3260

Ala Gly Ala Ile His Cys Arg Asn Phe Lys Cys Pro Ser Gly Thr Tyr 3265 3270 3275 3280

Cys Lys Asn Gly Asp Asn Gly Ser Ser Asn Cys Thr Glu Ile Thr Leu 3285 3290 3295

Gln Cys Pro Thr Asn Ser Gln Phe Thr Asp Cys Leu Pro Ser Cys Val 3300 3305 3310

Pro Ser Cys Ser Asn Arg Cys Glu Val Thr Ser Pro Ser Val Pro Ser 3315 3320 3325

Ser Cys Arg Glu Gly Cys Leu Cys Asn His Gly Phe Val Phe Ser Glu 3330 3335 3340

Asp Lys Cys Val Pro Arg Thr Gln Cys Gly Cys Lys Asp Ala Arg Gly 3345 3350 3355 3360

Ala Ile Ile Pro Ala Gly Lys Thr Trp Thr Ser Lys Gly Cys Thr Gln 3365 3370 3375

Ser Cys Ala Cys Val Glu Gly Asn Ile Gln Cys Gln Asn Phe Gln Cys 3380 3385 3390

Pro Pro Glu Thr Tyr Cys Lys Asp Asn Ser Glu Gly Ser Ser Thr Cys 3395 3400 3405

Thr Lys Ile Thr Leu Gln Cys Pro Ala His Thr Gln Tyr Thr Ser Cys 3410 3415 3420

Leu Pro Ser Cys Leu Pro Ser Cys Leu Asp Pro Glu Gly Leu Cys Lys 3425 3430 3435 3440

Asp Ile Ser Pro Lys Val Pro Ser Thr Cys Lys Glu Gly Cys Val Cys 3445 3450 3455

Gln Ser Gly Tyr Val Leu Asn Ser Asp Lys Cys Val Leu Arg Ala Glu 3460 3465 3470

Cys Asp Cys Lys Asp Ala Gln Gly Ala Leu Ile Pro Ala Gly Lys Thr Page 184

Trp Thr Ser Pro Gly Cys Thr Gln Ser Cys Ala Cys Met Gly Gly Ala 3490 3495 3500

Val Gln Cys Gln Ser Ser Gln Cys Pro Pro Gly Thr Tyr Cys Lys Asp 3505 3510 3515 3520

Asn Glu Asp Gly Asn Ser Asn Cys Ala Lys Ile Thr Leu Gln Cys Pro 3525 3530 3535

Ala His Ser Leu Phe Thr Asn Cys Leu Pro Ser Cys Leu Pro Ser Cys 3540 3550

Leu Asp Pro Asp Gly Leu Cys Lys Gly Ala Ser Pro Lys Val Pro Ser 3555 3560 3565

Thr Cys Lys Glu Gly Cys Ile Cys Gln Ser Gly Tyr Val Leu Ser Asn 3570 3575 3580

Asn Lys Cys Leu Leu Arg Asn Arg Cys Gly Cys Lys Asp Ala His Gly 3585 3590 3595 3600

Ala Leu Ile Pro Glu Asp Lys Thr Trp Val Ser Arg Gly Cys Thr Gln 3605 3610 3615

Ser Cys Val Cys Thr Gly Gly Ser Ile Gln Cys Leu Ser Phe Gln Cys 3620 3630

Pro Pro Gly Ala Tyr Cys Lys Asp Asn Glu Asp Gly Ser Ser Asn Cys 3635 3640 3645

Ala Arg Ile Pro Pro Gln Cys Pro Ala Asn Ser His Tyr Thr Asp Cys 3650 3655 3660

Phe Pro Pro Cys Pro Pro Ser Cys Ser Asp Pro Glu Gly His Cys Glu 3665 3670 3675 3680

Ala Ser Gly Pro Arg Val Pro Ser Thr Cys Arg Glu Gly Cys Leu Cys 3685 3690 3695

Asn Pro Gly Phe Val Leu Asp Arg Asp Lys Cys Val Pro Arg Val Glu 3700 3705 3710

Cys Gly Cys Lys Asp Ala Gln Gly Ala Leu Ile Pro Ser Gly Lys Thr 3715 3720 3725

Trp Thr Ser Pro Gly Arg Thr Gln Ser Cys Ala Cys Met Gly Gly Val 3730 3735 3740

Val Gln Cys Gln Ser Ser Gln Cys Pro Pro Gly Thr Tyr Cys Lys Asp 3745 3750 3755 3760

Asn Glu Asp Gly Asn Ser Asn Cys Ala Lys Ile Thr Leu Gln Cys Pro 3765 3770 3775

Thr His Ser Asn Tyr Thr Asp Cys Leu Pro Phe Cys Leu Pro Ser Cys 3780 3785 3790

Leu Asp Pro Ser Ala Leu Cys Gly Gly Thr Ser Pro Lys Gly Pro Ser 3795 3800 3805

Thr Cys Lys Glu Gly Cys Val Cys Gln Pro Gly Tyr Val Leu Asp Lys Page 185 Asp Lys Cys Ile Leu Lys Ile Glu Cys Gly Cys Lys Asp Thr Gln Gly 3825 3830 3835 3840

Ala Val Ile Pro Ala Gly Lys Thr Trp Leu Ser Thr Gly Cys Ile Gln 3845 3850 3855

Ser Cys Ala Cys Val Glu Gly Thr Ile Gln Cys Gln Asn Phe Gln Cys 3860 3865 3870

Pro Pro Gly Thr Tyr Cys Asn His Asn Asn Cys Ala Lys Ile Pro 3875 3880 3885

Leu Gln Cys Pro Ala His Ser His Phe Thr Ser Cys Leu Pro Ser Cys 3890 3895 3900

Pro Pro Ser Cys Ala Asn Leu Asp Gly Ser Cys Glu Gln Thr Ser Pro 3905 3910 3915 3920

Lys Val Pro Ser Thr Cys Lys Glu Gly Cys Leu Cys Gln Pro Gly Tyr 3925 3930 3935

Phe Leu Asn Asn Gly Lys Cys Val Leu Gln Thr His Cys Asp Cys Lys 3940 3945 3950

Asp Ala Glu Gly Gly Leu Val Pro Ala Gly Lys Thr Trp Thr Ser Lys 3955 3960 3965

Asp Cys Thr Gln Ser Cys Ala Cys Thr Gly Gly Ala Val Gln Cys Gln 3970 3975 3980

Asn Phe Gln Cys Pro Leu Gly Thr Tyr Cys Lys Asp Ser Gly Asp Gly 3985 3990 3995 4000

Ser Ser Asn Cys Thr Lys Ile His Lys Gly Ala Met Gly Asp Gly Val  $4005 \hspace{1cm} 4010 \hspace{1cm} 4015$ 

Leu Met Ala Gly Gly Ile Arg Ala Leu Gln Cys Pro Ala His Ser His 4020 4025 4030

Phe Thr Ser Cys Leu Pro Ser Cys Pro Pro Ser Cys Ser Asn Leu Asp 4035 4040 4045

Gly Ser Cys Val Glu Ser Asn Phe Lys Ala Pro Ser Val Cys Lys 4050 4055 4060

Gly Cys Ile Cys Gln Pro Gly Tyr Leu Leu Asn Asn Asp Lys Cys Val 4065 4070 4075 4080

Leu Arg Ile Gln Cys Gly Cys Lys Asp Thr Gln Gly Gly Leu Ile Pro 4085 4090 4095

Ala Gly Arg Thr Trp Ile Ser Ser Asp Cys Thr Lys Ser Cys Ser Cys 4100 4105 4110

Met Gly Gly Thr Ile Gln Cys Arg Asp Phe Gln Cys Pro Pro Gly Thr 4115 4120 4125

Tyr Cys Lys Glu Ser Asn Asp Ser Ser Arg Thr Cys Ala Lys Ile Pro 4130 4135 4140

Leu Gln Cys Pro Ala His Ser His Tyr Thr Asn Cys Leu Pro Ala Cys Page 186 Ser Arg Ser Cys Thr Asp Leu Asp Gly His Cys Glu Gly Thr Ser Pro 4165 4170 4175

Lys Val Pro Ser Pro Cys Lys Glu Gly Cys Leu Cys Gln Pro Gly Tyr 4180 4185 4190

Val Val His Asn His Lys Cys Val Leu Gln Ile His Cys Gly Cys Lys 4195 4200 4205

Asp Ala Gln Gly Gly Phe Val Pro Ala Gly Lys Thr Trp Ile Ser Arg 4210 4215 4220

Gly Cys Thr Gln Ser Cys Ala Cys Val Gly Gly Ala Val Gln Cys His 4225 4230 4235 4240

Asn Phe Thr Cys Pro Thr Gly Thr Gln Cys Gln Asn Ser Ser Cys Ser 4245 4250 4255

Lys Ile Thr Val Gln Cys Pro Ala His Ser His Tyr Thr Thr Cys Leu 4260 4265 4270

Pro Ser Cys Leu Pro Ser Cys Phe Asp Pro Glu Gly Leu Cys Gly Gly 4275 4280 4285

Ala Ser Pro Arg Ala Pro Ser Thr Cys Arg Glu Gly Cys Val Cys Glu 4290 4295 4300

Ala Asp Tyr Val Leu Arg Glu Asp Lys Cys Val Leu Arg Thr Gln Cys 4305 4310 4315 4320

Gly Cys Lys Asp Ala Gln Gly Asp Leu Ile Pro Ala Asn Lys Thr Trp 4325 4330 4335

Leu Thr Arg Gly Cys Ala Gln Lys Cys Thr Cys Lys Gly Gly Asn Ile 4340 4345 4350

His Cys Trp Asn Phe Lys Cys Pro Leu Gly Thr Glu Cys Lys Asp Ser 4355 4360 4365

Val Asp Gly Gly Ser Asn Cys Thr Lys Ile Ala Leu Gln Cys Pro Ala 4370 4375 4380

His Ser His His Thr Tyr Cys Leu Pro Ser Cys Ile Pro Ser Cys Ser 4385 4390 4395 4400

Asn Val Asn Asp Arg Cys Glu Ser Thr Ser Leu Gln Arg Pro Ser Thr 4405 4410 4415

Cys Ile Glu Gly Cys Leu Cys His Ser Gly Phe Val Phe Ser Lys Asp 4420 4425 4430

Lys Cys Val Pro Arg Thr Gln Cys Gly Cys Lys Asp Ser Gln Gly Thr 4435 4440 4445

Leu Ile Pro Ala Gly Lys Asn Trp Ile Thr Thr Gly Cys Ser Gln Arg 4450 4455 4460

Cys Thr Cys Thr Gly Gly Leu Val Gln Cys His Asp Phe Glu Cys Pro 4465 4470 4475 4480

Ser Gly Ala Glu Cys Gln Asp Ile Glu Asp Gly Asn Ser Asn Cys Val Page 187 Glu Ile Thr Val Gln Cys Pro Ala His Ser His Tyr Ser Lys Cys Leu 4500 4505 4510 4500 4505 Pro Pro Cys Gln Pro Ser Cys Ser Asp Pro Asp Gly His Cys Glu Gly 4515 4520 4525 Thr Ser Pro Glu Ala Pro Ser Thr Cys Glu Glu Gly Cys Val Cys Glu 4530 4535 4540 Pro Asp Tyr Val Leu Ser Asn Asp Lys Cys Val Pro Ser Ser Glu Cys Gly Cys Lys Asp Ala His Gly Val Leu Ile Pro Glu Ser Lys Thr Trp 4565 4570 4575 Val Ser Arg Gly Cys Thr Lys Asn Cys Thr Cys Lys Gly Gly Thr Val 4580 4585 4590 Gln Cys His Asp Phe Ser Cys Pro Thr Gly Ser Arg Cys Leu Asp Asn 4595 4600 4605 Asn Glu Gly Asn Ser Asn Cys Val Thr Tyr Ala Leu Lys Cys Pro Ala 4610 4615 4620 His Ser Leu Tyr Thr Asn Cys Leu Pro Ser Cys Leu Pro Ser Cys Ser 4625 4630 4635 4640 Asp Pro Glu Gly Leu Cys Gly Gly Thr Ser Pro Glu Val Pro Ser Thr Cys Lys Glu Gly Cys Ile Cys Gln Ser Gly Tyr Val Leu His Lys Asn 4660 4665 4670 Lys Cys Met Leu Arg Ile His Cys Asp Cys Lys Asp Phe Gln Gly Ser 4675 4680 4685 Leu Ile Lys Thr Gly Gln Thr Trp Ile Ser Ser Gly Cys Ser Lys Ile Cys Thr Cys Lys Gly Gly Phe Phe Gln Cys Gln Ser Tyr Lys Cys Pro 4705 4710 4715 4720 Ser Gly Thr Gln Cys Glu Glu Ser Glu Asp Gly Ser Ser Asn Cys Val 4725 4730 4735 Ser Ser Thr Met Lys Cys Pro Ala Asn Ser Leu Tyr Thr His Cys Leu Pro Thr Cys Leu Pro Ser Cys Ser Asn Pro Asp Gly Arg Cys Glu Gly 4755 4760 4765 Thr Ser His Lys Ala Pro Ser Thr Cys Arg Glu Gly Cys Val Cys Gln 4770 4780 Pro Gly Tyr Leu Leu Asn Lys Asp Thr Cys Val His Lys Asn Gln Cys 4785 4790 4795 4800 Gly Cys Lys Asp Ile Arg Gly Asn Ile Ile Pro Ala Gly Asn Thr Trp

Ile Ser Ser Asp Cys Thr Gln Ser Cys Ala Cys Thr Asp Gly Val Ile

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Gln Cys Gln Asn Phe Val Cys Pro Ser Gly Ser His Cys Gln Tyr Asn 4835 4840 4845

Glu Asp Gly Ser Ser Asp Cys Ala Ala Asn Lys Leu Glu Arg Cys Thr 4850 4855 4860

The Phe Gly Asp Pro Tyr Tyr Leu Thr Phe Asp Gly Phe Thr Tyr His 4865 4870 4875 4880

Phe Leu Gly Arg Met Asn Tyr Tyr Leu Ile Lys Thr Val Asp Lys Leu 4885 4890 4895

Pro Arg Gly Ile Glu Pro Leu Ile Met Glu Gly Arg Asn Lys Ile Ser 4900 4905 4910

Pro Lys Gly Ser Ser Thr Leu His Glu Val Thr Thr Ile Val Tyr Gly 4915 4920 4925

Tyr Lys Ile Gln Leu Gln Glu Glu Leu Val Val Leu Val Asn Asp Glu 4930 4935 4940

Lys Val Ala Val Pro Tyr Asn Pro Asn Glu His Leu Arg Val Met Leu 4945 4950 4955 4960

Arg Ala Gln Arg Leu Leu Leu Val Thr Asp Phe Glu Met Val Leu Asp 4965 4970 4975

Phe Asp Gly Lys His Ser Ala Val Ile Ser Leu Pro Thr Thr Tyr Arg 4980 4985 4990

Gly Leu Thr Arg Gly Leu Cys Gly Asn Tyr Asp Arg Asp Gln Ser Asn 4995 5000 5005

Glu Leu Met Leu Pro Ser Gly Val Leu Thr Ser Asn Val His Val Phe 5010 5020

Gly Asn Ser Trp Glu Val Lys Ala Gln His Ala Phe Phe Arg Phe Pro 5025 5030 5035 5040

Arg Ala Leu Pro Glu Asp Glu Glu Arg Asp Glu Glu Pro Asp Leu Leu 5045 5050 5055

Gln Ser Glu Cys Ser Gln Glu Gln Thr Ala Leu Ile Ser Ser Thr Gln 5060 5065 5070

Ala Cys Arg Val Leu Val Asp Pro Gln Gly Pro Phe Ala Ala Cys His 5075 5080 5085

Gln Ile Ile Ala Pro Glu Pro Phe Glu Gln Arg Cys Met Leu Asp Met 5090 5095 5100

Cys Thr Gly Trp Lys Thr Lys Glu Glu Glu Glu Leu Arg Cys Arg Val 5105 5110 5115 5120

Leu Ser Gly Tyr Ala Ile Ile Cys Gln Glu Ala Gly Ala Asn Met Thr 5125 5130 5135

Gly Trp Arg Asp His Thr His Cys Ala Met Thr Cys Pro Ala Asn Thr 5140 5145 5150

Val Tyr Gln Arg Cys Met Thr Pro Cys Pro Ala Ser Cys Ala Lys Phe Page 189

Val Thr Pro Lys Val Cys Glu Gly Pro Cys Val Glu Gly Cys Ala Ser 5170 5180

Leu Pro Gly Tyr Ile Tyr Ser Asp Thr Gln Ser Leu Pro Val Thr His 5185 5190 5195 5200

Cys Gly Cys Thr Ala Asp Gly Ile Tyr Tyr Lys Leu Gly Asp Ser Phe 5205 5210 5215

Val Thr Asn Asp Cys Ser Gln His Cys Thr Cys Ala Ser Gln Gly Ile 5220 5230

Leu Leu Cys Glu Pro Tyr Gly Cys Arg Ala Gly Glu Ser Cys Met Val 5235 5240 5245

Ala Asn Phe Thr Arg Gly Cys Phe Gln Asp Ser Pro Cys Leu Gln Asn 5250 5255 5260

Pro Cys His Asn Asp Gly Arg Cys Glu Glu Gln Gly Ala Thr Phe Ile 5265 5270 5275 5280

Cys His Cys Asp Phe Gly Tyr Gly Glu Phe Cys Thr Glu Pro Gln 5285 5290 5295

Asp Ile Thr Thr Arg Lys Lys Ile Glu Ala Ser Ser Leu Val Ala Ile 5300 5305 5310

Leu Pro Gly Val Leu Val Met Val Leu Val Pro Val Leu Leu Pro Arg 5315 5320 5325

Val Tyr Val Tyr Met Ala Thr Arg Thr Thr Met Gly Arg Arg Met 5330 5340

Lys Arg Lys Glu Lys Lys Leu Leu Arg Gln Ser Arg Leu Arg Leu Glu 5345 5350 5360

Asp Ala Asp Val Pro Glu Pro Thr Phe Lys Ala Thr Glu Phe

<210> 76 <211> 2601

<212> PRT

<213> Homo sapiens

<400> 76

Met Val Pro Pro Val Trp Thr Leu Leu Leu Leu Val Gly Ala Ala Leu 1 10 15

Phe Arg Lys Glu Lys Pro Pro Asp Gln Lys Leu Val Val Arg Ser Ser 20 25 30

Arg Asp Asn Tyr Val Leu Thr Gln Cys Asp Phe Glu Asp Asp Ala Lys 40 45

Pro Leu Cys Asp Trp Ser Gln Val Ser Ala Asp Asp Glu Asp Trp Val 50 60

Arg Ala Ser Gly Pro Ser Pro Thr Gly Ser Thr Gly Ala Pro Gly Gly 65 70 75 80

CURA2221.APP Tyr Pro Asn Gly Glu Gly Ser Tyr Leu His Met Glu Ser Asn Ser Phe 85 90 95 His Arg Gly Gly Val Ala Arg Leu Leu Ser Pro Asp Leu Trp Glu Gln
100 105 110 Gly Pro Leu Cys Val His Phe Ala His His Met Phe Gly Leu Ser Trp 115 120 125 Gly Ala Gln Leu Arg Leu Leu Leu Ser Gly Glu Glu Gly Arg Arg 130 135 140 Pro Asp Val Leu Trp Lys His Trp Asn Thr Gln Arg Pro Ser Trp Met 145 150 155 160 Leu Thr Thr Val Thr Val Pro Ala Gly Phe Thr Leu Pro Thr Arg Leu 165 170 175 Met Phe Glu Gly Thr Arg Gly Ser Thr Ala Tyr Leu Asp Ile Ala Leu 180 185 190 Asp Ala Leu Ser Ile Arg Arg Gly Ser Cys Asn Arg Val Cys Met Met 195 200 205 Gln Thr Cys Ser Phe Asp Ile Pro Asn Asp Leu Cys Asp Trp Thr Trp 210 220 Ile Pro Thr Ala Ser Gly Ala Lys Trp Thr Gln Lys Lys Gly Ser Ser 225 235 240 Gly Lys Pro Gly Val Gly Pro Asp Gly Asp Phe Ser Ser Pro Gly Ser 245 250 255 Gly Cys Tyr Met Leu Leu Asp Pro Lys Asn Ala Arg Pro Gly Gln Lys 260 265 270Ala Val Leu Leu Ser Pro Val Ser Leu Ser Ser Gly Cys Leu Ser Phe 275 280 285 Ser Phe His Tyr Ile Leu Arg Gly Gln Ser Pro Gly Ala Ala Leu His 290 295 300 Ile Tyr Ala Ser Val Leu Gly Ser Ile Arg Lys His Thr Leu Phe Ser 305 310 315 320Gly Gln Pro Gly Pro Asn Trp Gln Ala Val Ser Val Asn Tyr Thr Ala 325 330 335 Val Gly Arg Ile Gln Phe Ala Val Val Gly Val Phe Gly Lys Thr Pro 340 345 350 Glu Pro Ala Val Ala Val Asp Ala Thr Ser Ile Ala Pro Cys Gly Glu 355 360 365 Gly Phe Pro Gln Cys Asp Phe Glu Asp Asn Ala His Pro Phe Cys Asp 370 375 380 Trp Val Gln Thr Ser Gly Asp Gly Gly His Trp Ala Leu Gly His Lys 385 390 395 400 Asn Gly Pro Val His Gly Met Gly Pro Ala Gly Gly Phe Pro Asn Ala 405 410 415 CURA2221.APP
Gly Gly His Tyr Ile Tyr Leu Glu Ala Asp Glu Phe Ser Gln Ala Gly
420 425 Gln Ser Val Arg Leu Val Ser Arg Pro Phe Cys Ala Pro Gly Asp Ile 435 440 445 Cys Val Glu Phe Ala Tyr His Met Tyr Gly Leu Gly Glu Gly Thr Met 450 455 460 Leu Glu Leu Leu Gly Ser Pro Ala Gly Ser Pro Pro Ile Pro Leu 465 470 475 480 Trp Lys Arg Val Gly Ser Gln Arg Pro Tyr Trp Gln Asn Thr Ser Val 485 490 495 Thr Val Pro Ser Gly His Gln Gln Pro Met Gln Leu Ile Phe Lys Gly 500 505 510 Ile Gln Gly Ser Asn Thr Ala Ser Val Val Ala Met Gly Phe Ile Leu 515 520 525 Ile Asn Pro Gly Thr Cys Pro Val Lys Val Leu Pro Glu Leu Pro Pro 530 540 Val Ser Pro Val Ser Ser Thr Gly Pro Ser Glu Thr Thr Gly Leu Thr 545 550 560 Glu Asn Pro Thr Ile Ser Thr Lys Lys Pro Thr Val Ser Ile Glu Lys 565 570 575 Pro Ser Val Thr Thr Glu Lys Pro Thr Val Pro Lys Glu Lys Pro Thr 580 585 590 Ile Pro Thr Glu Lys Pro Thr Ile Ser Thr Glu Lys Pro Thr Ile Pro 595 600 605 Ser Glu Lys Pro Asn Met Pro Ser Glu Lys Pro Thr Ile Pro Ser Glu Lys Pro Thr Ile Leu Thr Glu Lys Pro Thr Ile Pro Ser Glu Lys Pro 625 630 635 640 Thr Ile Pro Ser Glu Lys Pro Thr Ile Ser Thr Glu Lys Pro Thr Val 645 650 655 Pro Thr Glu Glu Pro Thr Thr Pro Thr Glu Glu Thr Thr Ser Met 660 665 670 Glu Glu Pro Val Ile Pro Thr Glu Lys Pro Ser Ile Pro Thr Glu Lys 675 680 685 Pro Ser Ile Pro Thr Glu Lys Pro Thr Ile Ser Met Glu Glu Thr Ile 690 695 700 Ile Ser Thr Glu Lys Pro Thr Ile Ser Pro Glu Lys Pro Thr Ile Pro 705 710 715 720 Thr Glu Lys Pro Thr Ile Pro Thr Glu Lys Ser Thr Ile Ser Pro Glu 725 730 735 Lys Pro Thr Thr Pro Thr Glu Lys Pro Thr Ile Pro Thr Glu Lys Pro 740 745 750

CURA2221.APP Thr Ile Ser Pro Glu Lys Pro Thr Thr Pro Thr Glu Lys Pro Thr Ile 755 760 765 Ser Pro Glu Lys Leu Thr Ile Pro Thr Glu Lys Pro Thr Ile Pro Thr 770 775 780 Glu Lys Pro Thr Ile Pro Thr Glu Lys Pro Thr Ile Ser Thr Glu Glu 785 790 795 800 Pro Thr Thr Pro Thr Glu Glu Thr Thr Ile Ser Thr Glu Lys Pro Ser 805 810 815Ile Pro Met Glu Lys Pro Thr Leu Pro Thr Glu Glu Thr Thr Ser 820 825 830 Val Glu Glu Thr Thr Ile Ser Thr Glu Lys Leu Thr Ile Pro Met Glu 835 840 845 Pro Thr Ile Ser Thr Glu Lys Pro Thr Ile Pro Thr Glu Lys Pro 850 860 Thr Ile Ser Pro Glu Lys Leu Thr Ile Pro Thr Glu Lys Leu Thr Ile 865 870 875 880 Pro Thr Glu Lys Pro Thr Ile Pro Ile Glu Glu Thr Thr Ile Ser Thr 885 890 895 Glu Lys Leu Thr Ile Pro Thr Glu Lys Pro Thr Ile Ser Pro Glu Lys 900 905 910 Pro Thr Ile Ser Thr Glu Lys Pro Thr Ile Pro Thr Glu Lys Pro Thr 915 920 925 Ile Pro Thr Glu Glu Thr Thr Ile Ser Thr Glu Lys Leu Thr Ile Pro 930 935 940 Thr Glu Lys Pro Thr Ile Ser Pro Glu Lys Leu Thr Ile Pro Thr Glu 945 955 960 Lys Pro Thr Ile Ser Thr Glu Lys Pro Thr Ile Pro Thr Glu Lys Leu 965 970 975 Thr Ile Pro Thr Glu Lys Pro Thr Ile Pro Thr Glu Lys Pro Thr Ile 980 985 990 Pro Thr Glu Lys Leu Thr Ala Leu Arg Pro Pro His Pro Ser Pro Thr 1000 Ala Thr Gly Leu Ala Ala Leu Val Met Ser Pro His Ala Pro Ser Thr 1010 1015 1020 Pro Met Thr Ser Val Ile Leu Gly Thr Thr Thr Ser Arg Ser Ser 1025 1030 1035 Thr Glu Arg Cys Pro Pro Asn Ala Arg Tyr Glu Ser Cys Ala Cys Pro 1045 1050 1055 Ala Ser Cys Lys Ser Pro Arg Pro Ser Cys Gly Pro Leu Cys Arg Glu 1060 1065 1070 Gly Cys Val Cys Asn Pro Gly Phe Leu Phe Ser Asp Asn His Cys Ile 1075 1080 1085

Gln Ala Ser Ser Cys Asn Cys Phe Tyr Asn Asn Asp Tyr Tyr Glu Pro 1090 1095 1100

Gly Ala Glu Trp Phe Ser Pro Asn Cys Thr Glu His Cys Arg Cys Trp 1105 1110 1115 1120

Pro Gly Ser Arg Val Glu Cys Gln Ile Ser Gln Cys Gly Thr His Thr 1125 1130 1135

Val Cys Gln Leu Lys Asn Gly Gln Tyr Gly Cys His Pro Tyr Ala Gly 1140 1145 1150

Thr Ala Thr Cys Leu Val Tyr Gly Asp Pro His Tyr Val Thr Phe Asp 1155 1160 1165

Gly Arg His Phe Gly Phe Met Gly Lys Cys Thr Tyr Ile Leu Ala Gln 1170 1175 1180

Pro Cys Gly Asn Ser Thr Asp Pro Phe Phe Arg Val Thr Ala Lys Asn 1185 1190 1195 1200

Glu Glu Gln Gly Gln Glu Gly Val Ser Cys Leu Ser Lys Val 1205 1210 1215

Thr Leu Pro Glu Ser Thr Val Thr Leu Leu Lys Gly Arg Arg Thr Leu 1220 1225 1230

Val Gly Gly Gln Gln Val Thr Leu Pro Ala Ile Pro Ser Lys Gly Val 1235 1240 1245

Phe Leu Gly Ala Ser Gly Arg Phe Val Glu Leu Gln Thr Glu Phe Gly 1250 1260

Leu Arg Val Arg Trp Asp Gly Asp Gln Gln Leu Tyr Val Thr Val Ser 1265 1270 1275 1280

Ser Thr Tyr Ser Gly Lys Leu Cys Gly Leu Cys Gly Asn Tyr Asp Gly 1285 1290 1295

Asn Ser Asp Asn Asp His Leu Lys Leu Asp Gly Ser Pro Ala Gly Asp 1300 1305 1310

Lys Glu Glu Leu Gly Asn Ser Trp Gln Thr Asp Gln Asp Glu Asp Gln 1315 1320 1325

Glu Cys Gln Lys Tyr Gln Val Val Asn Ser Pro Ser Cys Asp Ser Ser 1330 1335 1340

Leu Gln Ser Ser Met Ser Gly Pro Gly Phe Cys Gly Arg Leu Val Asp 1345 1350 1355 1360

Thr His Gly Pro Phe Glu Thr Cys Leu Leu His Val Lys Ala Ala Ser 1365 1370 1375

Phe Phe Asp Ser Cys Met Leu Asp Met Cys Gly Phe Gln Gly Leu Gln 1380 1385 1390

His Leu Leu Cys Thr His Met Ser Thr Met Thr Thr Cys Gln Asp 1395 1400 1405

Ala Gly His Ala Val Lys Pro Trp Arg Glu Pro His Phe Cys Pro Met 1410 1415 1420

- Ala Cys Pro Pro Asn Ser Lys Tyr Ser Leu Cys Ala Lys Pro Cys Pro 1425 1430 1435 1440
- Asp Thr Cys His Ser Gly Phe Ser Gly Met Phe Cys Ser Asp Arg Cys  $1445 \hspace{1cm} 1450 \hspace{1cm} 1455$
- Val Glu Ala Cys Glu Cys Asn Pro Gly Phe Val Leu Ser Gly Leu Glu 1460 1465 1470
- Cys Ile Pro Arg Ser Gln Cys Gly Cys Leu His Pro Ala Gly Ser Tyr 1475 1480 1485
- Phe Lys Val Gly Glu Arg Trp Tyr Lys Pro Gly Cys Lys Glu Leu Cys 1490 1500
- Val Cys Glu Ser Asn Asn Arg Ile Arg Cys Gln Pro Trp Arg Cys Arg 1505 1510 1515 1520
- Ala Gln Glu Phe Cys Gly Gln Gln Asp Gly Ile Tyr Gly Cys His Ala 1525 1530 1535
- Gln Gly Ala Ala Thr Cys Thr Ala Ser Gly Asp Pro His Tyr Leu Thr 1540 1545 1550
- Phe Asp Gly Ala Leu His His Phe Met Gly Thr Cys Thr Tyr Val Leu 1555 1560 1565
- Thr Arg Pro Cys Trp Ser Arg Ser Gln Asp Ser Tyr Phe Val Val Ser 1570 1575 1580
- Ala Thr Asn Glu Asn Arg Gly Gly Ile Leu Glu Val Ser Tyr Ile Lys 1585 1590 1595 1600
- Ala Val His Val Thr Val Phe Asp Leu Ser Ile Ser Leu Leu Arg Gly 1605 1610 1615
- Cys Lys Val Met Leu Asn Gly His Arg Val Ala Leu Pro Val Trp Leu 1620 1625 1630
- Ala Gln Gly Arg Val Thr Ile Arg Leu Ser Ser Asn Leu Val Leu Leu 1635 1640 1645
- Tyr Thr Asn Phe Gly Leu Gln Val Arg Tyr Asp Gly Ser His Leu Val 1650 1660
- Glu Val Thr Val Pro Ser Ser Tyr Gly Gly Gln Leu Cys Gly Leu Cys 1665 1670 1680
- Gly Asn Tyr Asn Asn Asn Ser Leu Asp Asp Asn Leu Arg Pro Asp Arg 1685 1690 1695
- Lys Leu Ala Gly Asp Ser Met Gln Leu Gly Ala Ala Trp Lys Leu Pro 1700 1705 1710
- Glu Ser Ser Glu Pro Gly Cys Phe Leu Val Gly Gly Lys Pro Ser Ser 1715 1720 1725
- Cys Gln Glu Asn Ser Met Ala Asp Ala Trp Asn Lys Asn Cys Ala Ile 1730 1735 1740
- Leu Ile Asn Pro Gln Gly Pro Phe Ser Gln Cys His Gln Val Val Pro 1745 1750 1755 1760

Pro Gln Ser Ser Phe Ala Ser Cys Val His Gly Gln Cys Gly Thr Lys 1765 1770 1775

Gly Asp Thr Thr Ala Leu Cys Arg Ser Leu Gln Ala Tyr Ala Ser Leu 1780 1785 1790

Cys Ala Gln Ala Gly Gln Ala Pro Ala Trp Arg Asn Arg Thr Phe Cys 1795 1800 1805

Pro Met Arg Cys Pro Pro Gly Ser Ser Tyr Ser Pro Cys Ser Ser Pro 1810 1815 1820

Cys Pro Asp Thr Cys Ser Ser Ile Asn Asn Pro Arg Asp Cys Pro Lys 1825 1830 1835 1840

Ala Leu Pro Cys Ala Glu Ser Cys Glu Cys Gln Lys Gly His Ile Leu 1845 1850 1855

Ser Gly Thr Ser Cys Val Pro Leu Gly Gln Cys Gly Cys Thr Asp Pro 1860 1865 1870

Ala Gly Ser Tyr His Pro Val Gly Glu Arg Trp Tyr Thr Glu Asn Thr 1875 1880 1885

Cys Thr Arg Leu Cys Thr Cys Ser Val His Asn Asn Ile Thr Cys Phe 1890 1895 1900

Gln Ser Thr Cys Lys Pro Asn Gln Ile Cys Trp Ala Leu Asp Gly Leu 1905 1910 1915 1920

Leu Arg Cys Arg Ala Ser Gly Val Gly Val Cys Gln Leu Pro Gly Glu 1925 1930 1935

Ser His Tyr Val Ser Phe Asp Gly Ser Asn His Ser Ile Pro Asp Ala 1940 1945 1950

Cys Thr Leu Val Leu Val Lys Val Cys His Pro Ala Met Ala Leu Pro 1955 1960 1965

Phe Phe Lys Ile Ser Ala Lys His Glu Lys Glu Glu Gly Gly Thr Glu 1970 1975 1980

Ala Phe Arg Leu His Glu Val Tyr Ile Asp Ile Tyr Asp Ala Gln Val 1985 1990 1995 2000

Thr Leu Gln Lys Gly His Arg Val Leu Ile Asn Ser Lys Gln Val Thr 2005 2010 2015

Leu Pro Ala Ile Ser Gln Ile Pro Gly Val Ser Val Lys Ser Ser Ser 2020 2025 2030

Ile Tyr Ser Ile Val Asn Ile Lys Ile Gly Val Gln Val Lys Phe Asp 2035 2040 2045

Gly Asn His Leu Leu Glu Ile Glu Ile Pro Thr Thr Tyr Gly Lys 2050 2055 2060

Val Cys Gly Met Cys Gly Asn Phe Asn Asp Glu Glu Glu Asp Glu Leu 2065 2070 2075 2080

Met Met Pro Ser Asp Glu Val Ala Asn Ser Asp Ser Glu Phe Val Asn 2085 2090 2095

Ser Trp Lys Asp Lys Asp Ile Asp Pro Ser Cys Gln Ser Leu Leu Val 2100 2105 2110

Asp Glu Gln Gln Ile Pro Ala Glu Gln Gln Glu Asn Pro Ser Gly Asn 2115 2120 2125

Cys Arg Ala Ala Asp Leu Arg Arg Ala Arg Glu Lys Cys Glu Ala Ala 2130 2135 2140

Leu Arg Ala Pro Val Trp Ala Gln Cys Ala Ser Arg Ile Asp Leu Thr 2145 2150 2155 2160

Pro Phe Leu Val Asp Cys Ala Asn Thr Leu Cys Glu Phe Gly Gly Leu 2165 2170 2175

Tyr Gln Ala Leu Cys Gln Ala Leu Gln Ala Phe Gly Ala Thr Cys Gln 2180 2185 2190

Ser Gln Gly Leu Lys Pro Pro Leu Trp Arg Asn Ser Ser Phe Cys Pro 2195 2200 2205

Leu Glu Cys Pro Ala Tyr Ser Ser Tyr Thr Asn Cys Leu Pro Ser Cys 2210 2215 2220

Ser Pro Ser Cys Trp Asp Leu Asp Gly Arg Cys Glu Gly Ala Lys Val 2225 2230 2235 2240

Pro Ser Ala Cys Ala Glu Gly Cys Ile Cys Gln Pro Gly Tyr Val Leu 2245 2250 2255

Ser Glu Asp Lys Cys Val Pro Arg Ser Gln Cys Gly Cys Lys Asp Ala 2260 2265 2270

His Gly Gly Ser Ile Pro Leu Gly Lys Ser Trp Val Ser Ser Gly Cys 2275 2280 2285

Thr Glu Lys Cys Val Cys Thr Gly Gly Ala Ile Gln Cys Gly Asp Phe 2290 2295 2300

Arg Cys Pro Ser Gly Ser His Cys Gln Leu Thr Ser Asp Asn Ser Asn 2305 2310 2315 2320

Ser Asn Cys Val Ser Asp Lys Ser Glu Gln Cys Ser Val Tyr Gly Asp 2325 2330 2335

Pro Arg Tyr Leu Thr Phe Asp Gly Phe Ser Tyr Arg Leu Gln Gly Arg 2340 2345 2350

Met Thr Tyr Val Leu Ile Lys Thr Val Asp Val Leu Pro Glu Gly Val 2355 2360 2365

Glu Pro Leu Leu Val Glu Gly Arg Asn Lys Met Asp Pro Pro Arg Ser 2370 2375 2380

Ser Ile Phe Leu Gln Glu Val Ile Thr Thr Val Tyr Gly Tyr Lys Val 2385 2390 2395 2400

Gln Leu Gln Ala Gly Leu Glu Leu Val Val Asn Asn Gln Lys Met Ala 2405 2410 2415

Val Pro Tyr Arg Pro Asn Glu His Leu Arg Val Thr Leu Trp Gly Gln 2420 2425 2430

Arg Leu Tyr Leu Val Thr Asp Phe Glu Leu Val Val Ser Phe Gly Gly 2435 2440 2445

Arg Lys Asn Ala Val Ile Ser Leu Pro Ser Met Tyr Glu Gly Leu Val 2450 2455 2460

Ser Gly Leu Cys Gly Asn Tyr Asp Lys Asn Arg Lys Asn Asp Met Met 2465 2470 2475 2480

Leu Pro Ser Gly Ala Leu Thr Gln Asn Leu Asn Thr Phe Gly Asn Ser 2485 2490 2495

Trp Glu Val Lys Thr Glu Asp Ala Leu Leu Arg Phe Pro Arg Ala Ile 2500 2505 2510

Pro Ala Glu Glu Glu Gly Gln Gly Ala Glu Leu Gly Leu Arg Thr Gly 2515 2520 2525

Leu Gln Val Ser Glu Cys Ser Pro Glu Gln Leu Ala Ser Asn Ser Thr 2530 2535 2540

Gln Ala Cys Arg Val Leu Ala Asp Pro Gln Gly Pro Phe Ala Ala Cys 2545 2550 2555 2560

His Gln Thr Val Ala Pro Glu Pro Phe Gln Glu His Cys Val Leu Asp 2565 2570 2575

Leu Cys Ser Ala Gln Asp Pro Arg Glu Gln Glu Glu Leu Arg Cys Gln 2580 2585 2590

Val Leu Ser Gly Trp Ala Ala Ala Phe 2595 2600

<210> 77 <211> 170

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: MAM domain
 sequence

<400> 77

Cys Asp Phe Glu Asp Gly Ser His Pro Phe Cys Gly Trp Ser Gln Asp 1 5 10

Ser Gly Asp Asp Gly Asp Asp Leu Gln Trp Thr Arg Val Asn Ser Ala 20 25 30

Thr Gly Gly Ser Thr Gly Pro Arg Gly Asp His Thr Thr Gly Asn Gly 35 40 45

His Tyr Met Tyr Val Asp Thr Ser Ser Gly Leu Leu Gln Glu Gly Gln  $50 \ 55 \ 60$ 

Lys Ala Arg Leu Leu Ser Pro Pro Leu Pro Pro Asn Arg Ser Pro Glu 65 70 75 80

Cys Cys Leu Thr Phe Trp Tyr His Met Tyr Gly Ser Gly Val Gly Thr 85 90 95

Pro Gly Leu Asn Val Tyr Val Arg Glu Asn Gly Glu Thr Leu Leu Trp Page 198

Ser Arg Ser Gly His Gln Gly Gly Gln Trp Leu Leu Ala Glu Val Thr Leu Pro Thr Phe Ser Thr Lys Pro Phe Gln Val Val Phe Glu Gly Thr Arg Gly Gly Gly Ser Arg Gly Gly Ile Ala Leu Asp Asp Ile Ser Leu 160

105

Ser Thr His Ile Glu Gly Pro Cys Asn Gln 165 170

<210> 78

<211> 170

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: MAM domain sequence

Ser Gly Asp Asp Gly Asp Asp Leu Gln Trp Thr Arg Val Asn Ser Ala 20 25 30

Thr Gly Gly Ser Thr Gly Pro Arg Gly Asp His Thr Thr Gly Asn Gly 40 45

His Tyr Met Tyr Val Asp Thr Ser Ser Gly Leu Leu Gln Glu Gly Gln 50 60

Lys Ala Arg Leu Leu Ser Pro Pro Leu Pro Pro Asn Arg Ser Pro Glu 65 70 75 80

Cys Cys Leu Thr Phe Trp Tyr His Met Tyr Gly Ser Gly Val Gly Thr 85 90 95

Pro Gly Leu Asn Val Tyr Val Arg Glu Asn Gly Glu Thr Leu Leu Trp 100 105 110

Ser Arg Ser Gly His Gln Gly Gly Gln Trp Leu Leu Ala Glu Val Thr 115 120 125

Leu Pro Thr Phe Ser Thr Lys Pro Phe Gln Val Val Phe Glu Gly Thr 130 135 140

Arg Gly Gly Ser Arg Gly Gly Ile Ala Leu Asp Asp Ile Ser Leu 145 150 155 160

Ser Thr His Ile Glu Gly Pro Cys Asn Gln 165 170

<210> 79

<211> 812 <212> PRT

<213> Homo sapiens

Page 199

<400> 79 Met Gly Trp Arg Pro Arg Arg Ala Arg Gly Thr Pro Leu Leu Leu 1 10 15 Leu Leu Leu Leu Leu Trp Pro Val Pro Gly Ala Gly Val Leu Gln 20 25 30 Gly His Ile Pro Gly Gln Pro Val Thr Pro His Trp Val Leu Asp Gly 40 45Gln Pro Trp Arg Thr Val Ser Leu Glu Glu Pro Val Ser Lys Pro Asp 50 55 Met Gly Leu Val Ala Leu Glu Ala Glu Gly Gln Glu Leu Leu Leu Glu 65 70 75 80 Leu Glu Lys Asn His Arg Leu Leu Ala Pro Gly Tyr Ile Glu Thr His 85 90 95 Tyr Gly Pro Asp Gly Gln Pro Val Val Leu Ala Pro Asn His Thr Asp  $100 \hspace{1cm} 105 \hspace{1cm} 110$ His Cys His Tyr Gln Gly Arg Val Arg Gly Phe Pro Asp Ser Trp Val 115 120 125 Val Leu Cys Thr Cys Ser Gly Met Ser Gly Leu Ile Thr Leu Ser Arg 130 140 Asn Ala Ser Tyr Tyr Leu Arg Pro Trp Pro Pro Arg Gly Ser Lys Asp 145 150 155 160 Phe Ser Thr His Glu Ile Phe Arg Met Glu Gln Leu Leu Thr Trp Lys 165 170 175 Gly Thr Cys Gly His Arg Asp Pro Gly Asn Lys Ala Gly Met Thr Ser 180 185 Leu Pro Gly Gly Pro Gln Ser Arg Gly Arg Arg Glu Ala Arg Arg Thr 195 200 205 Arg Lys Tyr Leu Glu Leu Tyr Ile Val Ala Asp His Thr Leu Phe Leu 210 220 Thr Arg His Arg Asn Leu Asn His Thr Lys Gln Arg Leu Leu Glu Val 225 230 235 240 Ala Asn Tyr Val Asp Gln Leu Leu Arg Thr Leu Asp Ile Gln Val Ala 245 250 255 Leu Thr Gly Leu Glu Val Trp Thr Glu Arg Asp Arg Ser Arg Val Thr 260 265 270 Gln Asp Ala Asn Ala Thr Leu Trp Ala Phe Leu Gln Trp Arg Arg Gly 275 280 285 Leu Trp Ala Gln Arg Pro His Asp Ser Ala Gln Leu Leu Thr Gly Arg 290 295 300 Ala Phe Gln Gly Ala Thr Val Gly Leu Ala Pro Val Glu Gly Met Cys 305 310 315 320 Arg Ala Glu Ser Ser Gly Gly Val Ser Thr Asp His Ser Glu Leu Pro Page 200

Ile Gly Ala Ala Thr Met Ala His Glu Ile Gly His Ser Leu Gly  $340 \hspace{1cm} 345$ Leu Ser His Asp Pro Asp Gly Cys Cys Val Glu Ala Ala Ala Glu Ser 355 360 365 Gly Gly Cys Val Met Ala Ala Ala Thr Gly His Pro Phe Pro Arg Val 370 375 380 Phe Ser Ala Cys Ser Arg Arg Gln Leu Arg Ala Phe Phe Arg Lys Gly 385 390 400 Gly Gly Ala Cys Leu Ser Asn Ala Pro Asp Pro Gly Leu Pro Val Pro 405 410 415 Pro Ala Leu Cys Gly Asn Gly Phe Val Glu Ala Gly Glu Glu Cys Asp 420 425 430 Cys Gly Pro Gly Gln Glu Cys Arg Asp Leu Cys Cys Phe Ala His Asn 435 440 Cys Ser Leu Arg Pro Gly Ala Gln Cys Ala His Gly Asp Cys Cys Val 450 455 460 Arg Cys Leu Leu Lys Pro Ala Gly Ala Leu Cys Arg Gln Ala Met Gly 465 470 475 480 Asp Cys Asp Leu Pro Glu Phe Cys Thr Gly Thr Ser Ser His Cys Pro 485 490 495 Pro Asp Val Tyr Leu Leu Asp Gly Ser Pro Cys Ala Arg Gly Ser Gly 500 505 510 Tyr Cys Trp Asp Gly Ala Cys Pro Thr Leu Glu Gln Gln Cys Gln Gln 515 525 Leu Trp Gly Pro Gly Ser His Pro Ala Pro Glu Ala Cys Phe Gln Val 530 540 Val Asn Ser Ala Gly Asp Ala His Gly Asn Cys Gly Gln Asp Ser Glu 545 550 555 560 Gly His Phe Leu Pro Cys Ala Gly Arg Asp Ala Leu Cys Gly Lys Leu 565 570 575 Gln Cys Gln Gly Gly Lys Pro Ser Leu Leu Ala Pro His Met Val Pro 580 585 590 Val Asp Ser Thr Val His Leu Asp Gly Gln Glu Val Thr Cys Arg Gly 595 600 605 Ala Leu Ala Leu Pro Ser Ala Gln Leu Asp Leu Gly Leu Gly Leu 610 620 Val Glu Pro Gly Thr Gln Cys Gly Pro Arg Met Val Cys Gln Ser Arg 625 630 635 Arg Cys Arg Lys Asn Ala Phe Gln Glu Leu Gln Arg Cys Leu Thr Ala 645 650 655 Cys His Ser His Gly Val Cys Asn Ser Asn His Asn Cys His Cys Ala Page 201

<210> 80 <211> 728

<212> PRT

<213> Homo sapiens

CURA2221.APP Leu Asn His Thr Lys Gln Arg Leu Leu Glu Val Ala Asn Tyr Val Asp 145 150 155 Gln Leu Leu Arg Thr Leu Asp Ile Gln Val Ala Leu Thr Gly Leu Glu 165 170 175 Val Trp Thr Glu Arg Asp Arg Ser Arg Val Thr Gln Asp Ala Asn Ala 180 185 190 Thr Leu Trp Ala Phe Leu Gln Trp Arg Arg Gly Leu Trp Ala Gln Arg 195 200 205 Pro His Asp Ser Ala Gln Leu Leu Thr Gly Arg Ala Phe Gln Gly Ala 210 220 Thr Val Gly Leu Ala Pro Val Glu Gly Met Cys Arg Ala Glu Ser Ser 225 230 235 240 Gly Gly Val Ser Thr Asp His Ser Glu Leu Pro Ile Gly Ala Ala Ala 245 250 255 Thr Met Ala His Glu Ile Gly His Ser Leu Gly Leu Ser His Asp Pro 260 265 270 Asp Gly Cys Cys Val Glu Ala Ala Glu Ser Gly Gly Cys Val Met 275 280 285 Ala Ala Ala Thr Gly His Pro Phe Pro Arg Val Phe Ser Ala Cys Ser 290 295 300 Arg Arg Gln Leu Arg Ala Phe Phe Arg Lys Gly Gly Ala Cys Leu 305 310 315 320 Ser Asn Ala Pro Asp Pro Gly Leu Pro Val Pro Pro Ala Leu Cys Gly 325 330 335 Asn Gly Phe Val Glu Ala Gly Glu Glu Cys Asp Cys Gly Pro Gly Gln 340 345 350 Glu Cys Arg Asp Leu Cys Cys Phe Ala His Asn Cys Ser Leu Arg Pro 355 360 365 Gly Ala Gln Cys Ala His Gly Asp Cys Cys Val Arg Cys Leu Leu Lys 370 375 Pro Ala Gly Ala Leu Cys Arg Gln Ala Met Gly Asp Cys Asp Leu Pro 385 390 395 400 Glu Phe Cys Thr Gly Thr Ser Ser His Cys Pro Pro Asp Val Tyr Leu 405 410 415 Leu Asp Gly Ser Pro Cys Ala Arg Gly Ser Gly Tyr Cys Trp Asp Gly 420 425 430 Ala Cys Pro Thr Leu Glu Gln Gln Cys Gln Gln Leu Trp Gly Pro Gly 435 440 445 Ser His Pro Ala Pro Glu Ala Cys Phe Gln Val Val Asn Ser Ala Gly 450 460 Asp Ala His Gly Asn Cys Gly Gln Asp Ser Glu Gly His Phe Leu Pro 465 470 475 480

Cys Ala Gly Arg Asp Ala Leu Cys Gly Lys Leu Gln Cys Gln Gly Gly 485 490 495 Lys Pro Ser Leu Leu Ala Pro His Met Val Pro Val Asp Ser Thr Val 500 505 His Leu Asp Gly Gln Glu Val Thr Cys Arg Gly Ala Leu Ala Leu Pro 515 520 525 Ser Ala Gln Leu Asp Leu Leu Gly Leu Gly Leu Val Glu Pro Gly Thr 530 540 Gln Cys Gly Pro Arg Met Val Cys Gln Ser Arg Arg Cys Arg Lys Asn 545 550 560 Ala Phe Gln Glu Leu Gln Arg Cys Leu Thr Ala Cys His Ser His Gly
565 570 575 Val Cys Asn Ser Asn His Asn Cys His Cys Ala Pro Gly Trp Ala Pro 580 585 590 Pro Phe Cys Asp Lys Pro Gly Phe Gly Gly Ser Met Asp Ser Gly Pro 595 600 605 Val Gln Ala Glu Asn His Asp Thr Phe Leu Leu Ala Met Leu Leu Ser 610 615 620 Val Leu Leu Pro Leu Leu Pro Gly Ala Gly Leu Ala Trp Cys Cys Tyr 625 630 635 640 Arg Leu Pro Gly Ala His Leu Gln Arg Cys Ser Trp Gly Cys Arg Arg 655 655 Asp Pro Ala Cys Ser Gly Pro Lys Asp Gly Pro His Arg Asp His Pro  $660 \hspace{1.5cm} 665 \hspace{1.5cm} 670$ Leu Gly Gly Val His Pro Met Glu Leu Gly Pro Thr Ala Thr Gly Gln 675 680 685 Pro Trp Pro Leu Asp Pro Glu Asn Ser His Glu Pro Ser Ser His Pro 690 695 700 Glu Lys Pro Leu Pro Ala Val Ser Pro Asp Pro Gln Ala Asp Gln Val 705 710 715 720 Gln Met Pro Arg Ser Cys Leu Trp 725

<210> 81 <211> 802 <212> PRT

<213> Homo sapiens

Gln Pro Trp Arg Thr Val Ser Leu Glu Glu Pro Val Ser Lys Pro Asp 50 60 Met Gly Leu Val Ala Leu Glu Ala Glu Gly Gln Glu Leu Leu Leu Glu 65 70 75 80 Leu Glu Lys Asn His Arg Leu Leu Ala Pro Gly Tyr Ile Glu Thr His 85 90 95 Tyr Gly Pro Asp Gly Gln Pro Val Val Leu Ala Pro Asn His Thr Asp 100 105 110His Cys His Tyr Gln Gly Arg Val Arg Gly Phe Pro Asp Ser Trp Val 115 120 125 Val Leu Cys Thr Cys Ser Gly Met Ser Gly Leu Ile Thr Leu Ser Arg 130 135 140 Asn Ala Ser Tyr Tyr Leu Arg Pro Trp Pro Pro Arg Gly Ser Lys Asp 145 150 155 160 Phe Ser Thr His Glu Ile Phe Arg Met Glu Gln Leu Leu Thr Trp Lys 165 170 175 Gly Thr Cys Gly His Arg Asp Pro Gly Asn Lys Ala Gly Met Thr Ser 180 185 190 Leu Pro Gly Gly Pro Gln Ser Arg Gly Arg Arg Glu Ala Arg Arg Thr 195 200 205 Arg Lys Tyr Leu Glu Leu Tyr Ile Val Ala Asp His Thr Leu Phe Leu 210 215 220 Thr Arg His Arg Asn Leu Asn His Thr Lys Gln Arg Leu Leu Glu Val 225 235 240 Ala Asn Tyr Val **Asp Gln** Leu Leu Arg Thr Leu Asp Ile Gln Val Ala 245 250 255 Leu Thr Gly Leu Glu Val Trp Thr Glu Arg Asp Arg Ser Arg Val Thr 260 265 270 Gln Asp Ala Asn Ala Thr Leu Trp Ala Phe Leu Gln Trp Arg Arg Gly 275 280 285 Leu Trp Ala Gln Arg Pro His Asp Ser Ala Gln Leu Leu Thr Gly Arg 290 295 300 Ala Phe Gln Gly Ala Thr Val Gly Leu Ala Pro Val Glu Gly Met Cys 315 320 Arg Ala Glu Ser Ser Gly Gly Val Ser Thr Asp His Ser Glu Leu Pro 325 330 335 Ile Gly Ala Ala Ala Thr Met Ala His Glu Ile Gly His Ser Leu Gly 340 345 350 Leu Ser His Asp Pro Asp Gly Cys Cys Val Glu Ala Ala Glu Ser 355 360 365 Gly Gly Cys Val Met Ala Ala Ala Thr Gly His Pro Phe Pro Arg Val 370 380 Page 205

Phe Ser Ala Cys Ser Arg Arg Gln Leu Arg Ala Phe Phe Arg Lys Gly 385 390 400 Gly Gly Ala Cys Leu Ser Asn Ala Pro Asp Pro Gly Leu Pro Val Pro 405 410 415 Pro Ala Leu Cys Gly Asn Gly Phe Val Glu Ala Gly Glu Glu Cys Asp 420 425 430 Cys Gly Pro Gly Gln Glu Cys Arg Asp Leu Cys Cys Phe Ala His Asn 435 440 445 Cys Ser Leu Arg Pro Gly Ala Gln Cys Ala His Gly Asp Cys Cys Val 450 455 460 Arg Cys Leu Leu Lys Pro Ala Gly Ala Leu Cys Arg Gln Ala Met Gly 465 470 475 480 Asp Cys Asp Leu Pro Glu Phe Cys Thr Gly Thr Ser Ser His Cys Pro 485 490 495 Pro Asp Val Tyr Leu Leu Asp Gly Ser Pro Cys Ala Arg Gly Ser Gly 500 505 510 Tyr Cys Trp Asp Gly Ala Cys Pro Thr Leu Glu Gln Gln Cys Gln Gln 515 520 525 Leu Trp Gly Pro Gly Ser His Pro Ala Pro Glu Ala Cys Phe Gln Val 530 540 val Asn Ser Ala Gly Asp Ala His Gly Asn Cys Gly Gln Asp Ser Glu 545 550 555 Gly His Phe Leu Pro Cys Ala Gly Arg Asp Ala Leu Cys Gly Lys Leu 565 570 575 Gln Cys Gln Gly Gly Lys Pro Ser Leu Leu Ala Pro His Met Val Pro 580 585 590 Val Asp Ser Thr Val His Leu Asp Gly Gln Glu Val Thr Cys Arg Gly 595 600 605 Ala Leu Ala Leu Pro Ser Ala Gln Leu Asp Leu Leu Gly Leu Gly Leu 610 620 Val Glu Pro Gly Thr Gln Cys Gly Pro Arg Met Val Cys Gln Ser Arg 625 630 635 640 Arg Cys Arg Lys Asn Ala Phe Gln Glu Leu Gln Arg Cys Leu Thr Ala 645 650 655 Cys His Ser His Gly Ala Gly Leu His Pro Ser Val Thr Ser Gln Ala 660 670 Leu Val Ala Ala Trp Thr Val Ala Leu Cys Arg Leu Lys Thr Met Thr 675 680 685 Pro Ser Cys Trp Pro Cys Ser Ser Ala Ser Cys Cys Leu Cys Ser Gln 690 695 700 Gly Pro Ala Trp Pro Gly Val Ala Thr Asp Ser Gln Glu Pro Ile Cys 705 710 715 720 Page 206

Ser Asp Ala Ala Gly Ala Ala Glu Gly Thr Leu Arg Ala Val Ala Pro 735 Pro Lys Met Ala His Thr Gly Thr Thr Pro Trp Ala Ala Phe Thr Pro Trp Ser Trp Ala Pro Gln Pro Leu Asp Ser Pro Gly Pro Trp Thr Leu Arg 765 Thr Leu Arg 765 Thr Leu Arg Ser Leu Cys Gln Gln Ser Arg Leu Thr Pro Lys Ile Lys Ser Arg Cys Gln Asp Pro Ala Ser Gly 800 Glu Arg

<210> 82 <211> 685 <212> PRT

<213> Mus musculus

Adologo 82
Asp His Cys Gln Tyr His Gly Arg Val Arg Gly Phe Arg Glu Ser Trp
10
All Val Leu Ser Thr Cys Ser Gly Met Ser Gly Leu Ile Val Leu Ser
Ser Lys Val Ser Tyr Tyr Leu Gln Pro Arg Thr Pro Gly Asp Thr Lys
Asp Phe Pro Thr His Glu Ile Phe Arg Met Glu Gln Leu Phe Thr Trp
Arg Gly Val Gln Arg Asp Lys Asn Ser Gln Tyr Lys Ala Gly Met Ala
65
Ser Leu Pro His Val Pro Gln Ser Arg Val Arg Arg Glu Ala Arg Arg
Ser Pro Arg Tyr Leu Glu Leu Tyr Ile Val Ala Asp His Thr Leu Phe
110
Leu Leu Gln His Gln Asn Leu Asn His Thr Arg Gln Arg Leu Leu Glu
130
Ash Cys Val Asp Gln Ile Leu Arg Thr Leu Asp Ile Gln Leu
Val Ala Asn Cys Val Asp Gln Ile Leu Arg Thr Leu Asp Ile Gln Leu
Val Leu Thr Gly Leu Glu Val Trp Thr Glu Gln Asp Leu Ser Arg Ile
145
Gly Val Trp Ala Arg Arg Pro His Asp Ser Thr Gln Leu Glu Asp Met
Page 207

Pro Arg Gly Glu Leu Ser Phe Gly Gly Val Ser Thr Asp His Ser Glu 210 215 220 Leu Pro Ile Gly Thr Ala Ala Thr Met Ala His Glu Ile Gly His Ser 225 230 235 240 Leu Gly Leu His His Asp Pro Glu Gly Cys Cys Val Gln Ala Asp Ala 245 250 255 Glu Gln Gly Gly Cys Val Met Glu Ala Ala Thr Gly His Pro Phe Pro 260 265 270 Arg Val Phe Ser Ala Cys Ser Arg Arg Gln Leu Arg Thr Phe Phe Arg 275 280 285 Lys Gly Gly Gly Pro Cys Leu Ser Asn Thr Ser Ala Pro Gly Leu Leu 290 295 300 Val Leu Pro Ser Arg Cys Gly Asn Gly Phe Leu Glu Ala Gly Glu Glu 305 310 315 320 Cys Asp Cys Gly Ser Gly Gln Lys Cys Pro Asp Pro Cys Cys Phe Ala 325 330 335 His Asn Cys Ser Leu Arg Ala Gly Ala Gln Cys Ala His Gly Asp Cys 340 345 350 Cys Ala Arg Cys Leu Leu Lys Ser Ala Gly Thr Pro Cys Arg Pro Ala 355 360 365 Ala Thr Asp Cys Asp Leu Pro Glu Phe Cys Thr Gly Thr Ser Pro Tyr 370 375 380 Cys Pro Ala Asp Val Tyr Leu Leu Asp Gly Ser Pro Cys Ala Glu Gly 385 390 395 400 Arg Gly Tyr Cys Leu Asp Gly Trp Cys Pro Thr Leu Glu Gln Gln Cys 405 410 415Gln Gln Leu Trp Gly Pro Gly Ser Lys Pro Ala Pro Glu Pro Cys Phe  $420 \hspace{1cm} 425 \hspace{1cm} 430$ Gln Gln Met Asn Ser Met Gly Asn Ser Gln Gly Asn Cys Gly Gln Asp 435 440 445 His Lys Gly Ser Phe Leu Pro Cys Ala Gln Arg Asp Ala Leu Cys Gly 450 455 460 Lys Leu Leu Cys Gln Gly Gly Glu Pro Asn Pro Leu Val Pro His Ile 465 470 475 480 Val Thr Met Asp Ser Thr Ile Leu Leu Glu Gly Arg Glu Val Val Cys 485 490 495 Arg Gly Ala Phe Val Leu Pro Asp Ser His Leu Asp Gln Leu Asp Leu 500 510 Gly Leu Val Glu Pro Gly Thr Gly Cys Gly Pro Arg Met Val Cys Gln 515 520 Asp Arg His Cys Gln Asn Ala Thr Ser Gln Glu Leu Glu Arg Cys Leu Page 208

 Thr 545
 Ala Cys
 His Asn 550
 Gly 550
 Cys
 Asn 555
 Asn 555
 Asn Arg Asn Cys 560
 Cys 560

 Cys
 Ala Ala Ala Gly Trp 565
 Ala Pro Pro Pro Pro S70
 Asp Lys Pro Gly Leu Gly 575
 Gly 575

<210> 83 <211> 914 <212> PRT <213> Xenopus laevis

<400> 83 Met Gly Thr Glu Gly Arg Leu Ser Thr Trp Leu Gly Leu Gly Ala Val 1 1 1

Ile Val Gly Leu Leu Leu Pro Pro Val Leu Thr Leu Gly Ala His Gln 20 25 30

Gly Glu Leu Val Thr Ala Phe Trp Leu Gln Asn Gly Arg Ala Lys Arg
35 40 45

Ser Val Asp Leu Leu Asp Lys Gly Thr Pro Asp Gly Glu Ile Leu 50 60

Val Ser Ser Glu Gly Arg Lys Phe Ile Leu Lys Val Glu Arg Asn His 65 70 75 80

Leu Leu Phe Ala Pro Gly Tyr Thr Glu Thr His Tyr Thr Asp Gly Gln  $85 \hspace{1cm} 90 \hspace{1cm} 95$ 

Met Val Thr Leu Ser Pro Asn His Thr Glu His Cys Tyr Tyr His Gly  $100 \hspace{1cm} 105 \hspace{1cm} 110$ 

Gln Val Glu Asn Tyr Asp Glu Ser Ser Val Ala Leu Thr Thr Cys Ser 115 120 125

Gly Ile Ser Gly Leu Ile Trp Leu Ser Thr Asn Asn Ser Tyr Tyr Leu 130 135 140

CURA2221.APP Lys Pro Leu Glu Val Pro Gly Lys Glu Thr His Thr Leu Val Arg Thr 145 150 155 160 Glu His Leu Leu Ile Lys Glu Gly Ser Cys Gly His Asp Gly His Ser 165 170 175 Gly Ser Thr Ala Ser Tyr Leu Gln Glu Phe Thr Ala Pro Ser Ser His 180 185 190 His His Arg Val Arg Arg Asn Val Trp Arg Ser Gln Lys Tyr Met Glu 195 200 205Leu Phe Ile Val Ala Asp Tyr Ser Met Phe Met Lys Gln Asn Arg Asn 210 220 Leu Gly Ser Thr Lys Gln Arg Val Leu Glu Ile Ala Asn Tyr Val Asp 225 230 235 240 Lys Phe Tyr Met Ser Met Asn Ile Lys Val Ala Leu Ile Gly Leu Glu 245 250 255Val Trp Thr Glu Arg Asp Gln Cys Glu Val Asn Asp Asp Ala Asn Asp 260 265 270 Ser Leu Lys Ser Phe Leu Gln Trp Lys Gln Lys Leu Arg Ser Arg Lys 275 280 285 Lys His Asp Asn Ala Gln Leu Ile Thr Gly Val Thr Phe Lys Gly Thr 290 295 300 Thr Ile Gly Met Ala Pro Leu Glu Gly Met Cys Thr Ala Glu Asn Ser 305 310 315 320Gly Gly Val Ser Met Asp His Ser Glu Asn Ala Ile Gly Ala Ala Ala 325 330 335 Thr Met Ala His Glu Ile Gly His Asn Phe Gly Met Ser His Asp Asp 340 345 350 Gly Cys Cys Val Glu Ala Thr Pro Glu Gln Gly Gly Cys Ile Met Ala 355 360 365 Ala Ala Thr Gly His Pro Phe Pro Arg Lys Phe Ser Ser Cys Ser Gln 370 375 380Lys Gln Leu Met Ser Tyr Phe Gln Lys Gly Gly Gly Met Cys Leu Phe 385 390 395 400 Asn Met Pro Asn Thr Lys Asp Leu Val Met Gly Lys Lys Cys Gly Asn 405 410 415 Gly Phe Leu Glu Glu Glu Glu Gln Cys Asp Cys Gly Glu Pro Glu Glu 420 425 430 Cys Thr Asn Ser Cys Cys Asn Ala Asn Asn Cys Thr Leu Lys Ala Gly 435 440 445 Ala Gln Cys Ala His Gly Glu Cys Cys Gln Asp Cys Lys Leu Lys Ser 450 455 460 Ala Gly Thr Gln Cys Arg Glu Met Ala Gly Ser Cys Asp Leu Pro Glu 465 470 475 480

CURA2221.APP Phe Cys Thr Gly Asp Ala Pro Ser Cys Pro Ser Asn Val Tyr Lys Leu 485 490 495 Asp Gly Ser Leu Cys Ala Asp Gly Asn Ala Tyr Cys Tyr Asn Gly Met 500 505 510 Cys Leu Thr His Gln Gln Gln Cys Ile His Leu Trp Gly Ser Gly Ala 515 520 525 Val Val Ala Pro Asn Phe Cys Phe Gln Asp Val Asn Lys Ala Gly Asp 530 540 Gln Tyr Gly Asn Cys Gly Lys Asn Gly Arg Gly Gln Phe Val Lys Cys 545 550 560 Thr Ser Arg Asp Ala Lys Cys Gly Lys Ile Gln Cys Gln Thr Ser Ser 575 Glu Lys Pro Arg Asp Pro Ser Met Val Lys Val Asp Asn Thr Ile Ile 580 590 Ile Asn Gly Tyr Lys Met Lys Cys Gln Gly Val His Ala Tyr Ser Met 595 600 605 Gln Glu Glu Gly Asp Pro Gly Leu Val Met Thr Gly Thr Lys Cys 610 620 Gly Asp Gly Met Val Cys Lys Asp Arg Arg Cys Gln Asn Ala Ser Phe 625 630 635 640 Phe Glu Leu Asp Gln Cys Val Ser Lys Cys Asn Gly His Gly Val Cys 655 Asn Ser Asn Arg Asn Cys His Cys Asp Ser Gly Trp Ala Pro Pro Tyr 660 670 Cys Asp Lys Pro Gly Pro Gly Gly Ser Gln Asp Ser Gly Pro Ala Pro 675 680 685 Ser Asp Leu Pro Val Gly Val Thr Ile Phe Leu Val Ile Leu Phe Leu 690 695 700 Val Leu Leu Ala Leu Ala Phe Ala Met Val Tyr Trp Tyr Arg Lys 705 710 715 720 Pro Gly Ser Leu Leu Asn Arg Trp Leu Met Lys Ser Lys Ala Lys Cys 725 730 735 Ser Leu Cys Lys Ala Thr Gln Pro Lys Ala Asn Arg Ala Tyr Ser Ser 740 745 750 Arg Ile Phe Thr Leu Arg Asn Ile Ser Tyr Pro Val Lys Ser Thr Ser 765 Lys Glu Thr Arg Ser Arg Asp Ile Phe Gln Gly Lys Thr Thr Ala Ala 770 780 Gln Asn Ser Ser Gln Pro Val Asn Val Val Arg Pro Leu Arg Pro Ala 785 790 795 800 Pro Ser Pro Val Ile Gln His Gly Val Gln Val Lys Pro Leu Arg Pro 805 810 815

Pro Pro Pro Pro Pro Met Lys Pro Ser Pro Ile Leu Pro Ala Lys Glu Gln
Thr Val His Val Lys Leu Leu Pro Pro Lys Lys Pro Leu Pro Ser Cys
835 Pro Ile Arg Thr Gln Gln Leu Asn Pro Pro Ser Lys Pro Leu Pro Val
855 Pro Ala His Lys Glu Pro Leu Leu Val Leu Thr Pro Ala Thr His 865 Pro Pro Ile Thr Asn Ser Ala Thr Gln Leu Lys Gly Pro His Arg
886 Pro Ile Gln Gly Gly Lys Val Gln Ala Ala Ala Ala Ala Phe Leu Gln
Arg Lys

<210> 84 <211> 203 <212> PRT

<213> Artificial Sequence

4400> 84
Lys Tyr Ile Glu Leu Val Ile Val Val Asp His Gly Met Tyr Thr Lys
Tyr Gly Ser Asp Leu Asn Lys Ile Arg Gln Arg Val His Gln Ile Val
Asn Leu Val Asn Glu Ile Tyr Arg Pro Gln Leu Asn Ile Arg Val Val
Leu Val Gly Leu Glu Ile Trp Ser Asp Gly Asp Lys Ile Asn Val Gln
Ser Asp Ala Asn Asp Thr Leu His Ser Phe Gly Glu Trp Arg Glu Thr
65 Asp Leu Leu Lys Arg Lys Ser His Asp Asn Ala Gln Leu Leu Thr Gly
95
Ile Asp Phe Asp Gly Asn Thr Ile Gly Ala Ala Tyr Val Gly Gly Met
Cys Ser Pro Lys Arg Ser Val Gly Val Val Gln Asp His Ser Pro Ile
Val Leu Leu Val Ala Val Thr Met Ala His Glu Leu Gly His Asn Leu
Gly Met Thr His Asp Asp Lys Asn Lys Asp Gly Cys Thr Cys Glu Gly
160

Gly Gly Ser Cys Ile Met Asn Pro Val Ala Ser Ser Ser Pro Ser Lys

Page 212

Lys Lys Phe Ser Asn Cys Ser Lys Asp Asp Tyr Gln Lys Phe Leu Thr 180 185 190

Lys Gln Lys Pro Gln Cys Leu Leu Asn Lys Pro 195 200

<210> 85

<211> 119 <212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:
 Pep\_M12B\_Propep (Reprolysin family propeptide)
 domain sequence

Thr Tyr Asp Glu Asp Gly Thr Leu Val Thr Glu Glu Pro Leu Ile Gln
20 25 30

Asp Asp His Cys Tyr Tyr Gln Gly Tyr Val Glu Gly Tyr Pro Asn Ser 40 45

Ala Val Ser Leu Ser Thr Cys Ser Gly Gly Leu Arg Gly Ile Leu Gln 50 60

Leu Glu Asn Leu Ser Tyr Gly Ile Glu Pro Leu Glu Ser Ser Asp Gly 65 70 75 80

Phe Glu His Ile Ile Tyr Gln Ile Glu Asn Asp Lys Thr Glu Pro Ser  $85 \hspace{1cm} 90 \hspace{1cm} 95$ 

Pro Cys Gly Glu Cys Gly Ser Leu Ser Thr Ser Thr Asp Ser Ser Tyr 100 105 110

Gly Ile Arg Ser Ala Ser Pro 115

<210> 86

<211> 422

<212> PRT

<213> Homo sapiens

<400> 86

Met Phe Ser Asn Ser Asp Glu Ala Val Ile Asn Lys Lys Leu Pro Lys 1 5 10 15

Glu Leu Leu Arg Ile Phe Ser Phe Leu Asp Val Val Thr Leu Cys 20 25 30

Arg Cys Ala Gln Val Ser Arg Ala Trp Asn Val Leu Ala Leu Asp Gly
35 40 45

Ser Asn Trp Gln Arg Ile Asp Leu Phe Asp Phe Gln Arg Asp Ile Glu 50 60

Gly Arg Val Val Glu Asn Ile Ser Lys Arg Cys Gly Gly Phe Leu Arg
65 70 75 80 Lys Leu Ser Leu Arg Gly Cys Leu Gly Val Gly Asp Asn Ala Leu Arg 85 90 95 Thr Phe Ala Gln Asn Cys Arg Asn Ile Glu Val Leu Asn Leu Asn Gly 100 105 110Cys Thr Lys Thr Thr Asp Ala Thr Cys Thr Ser Leu Ser Lys Phe Cys 115 120 125Ser Lys Leu Arg His Leu Asp Leu Ala Ser Cys Thr Ser Ile Thr Asn 130 135 140 Met Ser Leu Lys Ala Leu Ser Glu Gly Cys Pro Leu Leu Glu Gln Leu 145 150 155 160 Asn Ile Ser Trp Cys Asp Gln Val Thr Lys Asp Gly Ile Gln Ala Leu 165 170 175 Val Arg Gly Cys Gly Gly Leu Lys Ala Leu Phe Leu Lys Gly Cys Thr 180 185 190 Gln Leu Glu Asp Glu Ala Leu Lys Tyr Ile Gly Ala His Cys Pro Glu 195 200 205 Leu Val Thr Leu Asn Leu Gln Thr Cys Leu Gln Ile Thr Asp Glu Gly 210 220 Leu Ile Thr Ile Cys Arg Gly Cys His Lys Leu Gln Ser Leu Cys Ala 225 230 235 240 Ser Gly Cys Ser Asn Ile Thr Asp Ala Ile Leu Asn Ala Leu Gly Gln 245 250 255 Asn Cys Pro Arg Leu Arg Ile Leu Glu Val Ala Arg Cys Ser Gln Leu 260 265 270 Thr Asp Val Gly Phe Thr Thr Leu Ala Arg Asn Cys His Glu Leu Glu 275 280 285 Lys Met Asp Leu Glu Glu Cys Val Gln Ile Thr Asp Ser Thr Leu Ile 290 295 300 Gln Leu Ser Ile His Cys Pro Arg Leu Gln Val Leu Ser Leu Ser His 305 310 315 320 Cys Glu Leu Ile Thr Asp Asp Gly Ile Arg His Leu Gly Asn Gly Ala 325 330 335 Cys Ala His Asp Gln Leu Glu Val Ile Glu Leu Asp Asn Cys Pro Leu 340 345 350Ile Thr Asp Ala Ser Leu Glu His Leu Lys Ser Cys His Ser Leu Glu 355 360 365 Arg Ile Glu Leu Tyr Asp Cys Gln Gln Ile Thr Arg Ala Gly Ile Lys 370 380 Arg Leu Arg Thr His Leu Pro Asn Ile Lys Val His Ala Tyr Phe Ala 385 390 395 400

Pro Val Thr Pro Pro Pro Ser Val Gly Gly Ser Arg Gln Arg Phe Cys 405

Arg Cys Cys Ile Ile Leu 420

<210> 87 <211> 422 <212> PRT <213> Mus musculus

<213> Mus musculus <400> 87 Met Phe Ser Asn Ser Asp Glu Ala Val Ile Asn Lys Lys Leu Pro Lys 1 5 10 15 Glu Leu Leu Arg Ile Phe Ser Phe Pro Asp Val Val Thr Leu Cys 20 25 30 Arg Cys Ala Gln Val Ser Arg Ala Trp Asn Val Leu Ala Leu Asp Gly 35 40 45 Ser Asn Trp Gln Arg Ile Asp Leu Phe Asp Phe Gln Arg Asp Ile Glu 50 60 Gly Arg Val Glu Asn Ile Ser Lys Arg Cys Gly Gly Phe Leu Arg 65 70 75 80 Lys Leu Ser Leu Arg Gly Cys Leu Gly Val Gly Asp Asn Ala Leu Arg 85 90 95 Thr Phe Ala Gln Asn Cys Arg Asn Ile Glu Val Leu Ser Leu Asn Gly  $100 \hspace{1cm} 105 \hspace{1cm} 110$ Cys Thr Lys Thr Thr Asp Ala Thr Cys Thr Ser Leu Ser Lys Phe Cys 115 120 125 Ser Lys Leu Arg His Leu Asp Leu Ala Ser Cys Thr Ser Ile Thr Asn 130 135 140 Met Ser Leu Lys Ala Leu Ser Glu Gly Cys Pro Leu Leu Glu Gln Leu 145 150 155 160 Asn Ile Ser Trp Cys Asp Gln Val Thr Lys Asp Gly Ile Gln Ala Leu 165 170 175 Val Arg Gly Cys Gly Gly Leu Lys Ala Leu Phe Leu Lys Gly Cys Thr 180 185 190 Gln Leu Glu Asp Glu Ala Leu Lys Tyr Ile Gly Ala His Cys Pro Glu 195 200 205 Leu Val Thr Leu Asn Leu Gln Thr Cys Leu Gln Ile Thr Asp Glu Gly 210 220 Leu Ile Thr Ile Cys Arg Gly Cys His Lys Leu Gln Ser Leu Cys Ala 225 230 240 Ser Gly Cys Ser Asn Ile Thr Asp Ala Ile Leu Asn Ala Leu Gly Gln 245 250 255 Asn Cys Pro Arg Leu Arg Ile Leu Glu Val Ala Arg Cys Ser Gln Leu 260 265 270 Page 215

Thr Asp Val Gly Phe Thr Thr Leu Ala Arg Asn Cys His Glu Leu Glu
Lys Met Asp Leu Glu Glu Cys Val Gln Ile Thr Asp Ser Thr Leu Ile
Gln Leu Ser Ile His Cys 310 Pro Arg Leu Gln Val Leu Ser Leu Ser His
305 Cys Glu Leu Ile Thr Asp Asp Gly Ile Arg His Leu Gly Asn Gly Ala
325
Cys Ala His Asp Gln Leu Glu Val Ile Glu Leu Asp Asn Cys Pro Leu
345
Cys Ala His Asp Ala Ser Leu Glu His Leu Lys Ser Cys Pro Ser Phe Glu
Arg Ile Glu Leu Tyr Asp Cys Gln Gln Ile Thr Arg Ala Gly Ile Lys
Arg Leu Arg Thr His Leu Pro Asn Ile Lys Val His Ala Tyr Phe Ala
400
Pro Val Thr Pro Pro Pro Pro Ser Val Gly Gly Ser Arg Gln Arg Phe Cys
Arg Cys Cys Ile Ile Leu

<210> 88

<211> 423

<212> PRT

<213> Homo sapiens

<400> 88

Met Val Phe Ser Asn Asn Asp Glu Gly Leu Ile Asn Lys Lys Leu Pro 1 5 10

Lys Glu Leu Leu Leu Arg Ile Phe Ser Phe Leu Asp Ile Val Thr Leu 20 25 30

Cys Arg Cys Ala Gln Ile Ser Lys Ala Trp Asn Ile Leu Ala Leu Asp 35 40 45

Gly Ser Asn Trp Gln Arg Ile Asp Leu Phe Asn Phe Gln Thr Asp Val
50 55 60

Glu Gly Arg Val Val Glu Asn Ile Ser Lys Arg Cys Gly Gly Phe Leu 65 70 75 80

Arg Lys Leu Ser Leu Arg Gly Cys Ile Gly Val Gly Asp Ser Ser Leu  $85 \hspace{1cm} 90 \hspace{1cm} 95$ 

Lys Thr Phe Ala Gln Asn Cys Arg Asn Ile Glu His Leu Asn Leu Asn  $100 \hspace{1.5cm} 105 \hspace{1.5cm} 110$ 

Gly Cys Thr Lys Ile Thr Asp Ser Thr Cys Tyr Ser Leu Ser Arg Phe 115 120 125

Cys Ser Lys Leu Lys His Leu Asp Leu Thr Ser Cys Val Ser Ile Thr Page 216

130 135

Asn Ser Ser Leu Lys Gly Ile Ser Glu Gly Cys Arg Asn Leu Glu Tyr 145 150 155 160 Leu Asn Leu Ser Trp Cys Asp Gln Ile Thr Lys Asp Gly Ile Glu Ala 165 170 175 Leu Val Arg Gly Cys Arg Gly Leu Lys Ala Leu Leu Leu Arg Gly Cys 180 185 190 Thr Gln Leu Glu Asp Glu Ala Leu Lys His Ile Gln Asn Tyr Cys His 195 200 205 Glu Leu Val Ser Leu Asn Leu Gln Ser Cys Ser Arg Ile Thr Asp Glu 210 215 220 Gly Val Val Gln Ile Cys Arg Gly Cys His Arg Leu Gln Ala Leu Cys 225 230 240 Leu Ser Gly Cys Ser Asn Leu Thr Asp Ala Ser Leu Thr Ala Leu Gly 245 250 255 Leu Asn Cys Pro Arg Leu Gln Ile Leu Glu Ala Ala Arg Cys Ser His 260 265 270 Leu Thr Asp Ala Gly Phe Thr Leu Leu Ala Arg Asn Cys His Glu Leu 275 280 285 Glu Lys Met Asp Leu Glu Glu Cys Ile Leu Ile Thr Asp Ser Thr Leu 290 295 300 Ile Gln Leu Ser Ile His Cys Pro Lys Leu Gln Ala Leu Ser Leu Ser 305 310 315 320 His Cys Glu Leu Ile Thr Asp Asp Gly Ile Leu His Leu Ser Asn Ser 325 330 335 Thr Cys Gly His Glu Arg Leu Arg Val Leu Glu Leu Asp Asn Cys Leu 340 350 Leu Ile Thr Asp Val Ala Leu Glu His Leu Glu Asn Cys Arg Gly Leu 355 360 365 Glu Arg Leu Glu Leu Tyr Asp Cys Gln Gln Val Thr Arg Ala Gly Ile 370 375 380 Lys Arg Met Arg Ala Gln Leu Pro His Val Lys Val His Ala Tyr Phe 385 390 395 400 Ala Pro Val Thr Pro Pro Thr Ala Val Ala Gly Ser Gly Gln Arg Leu 405 415 Cys Arg Cys Cys Val Ile Leu 420

<sup>&</sup>lt;210> 89 <211> 425

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Homo sapiens

<sup>&</sup>lt;400> 89

CURA2221.APP Ser Ala Met Val Phe Ser Asn Asn Asp Glu Gly Leu Ile Asn Lys Lys 1 10 15 Leu Pro Lys Glu Leu Leu Leu Arg Ile Phe Ser Phe Leu Asp Ile Val 20 25 30 Thr Leu Cys Arg Cys Ala Gln Ile Ser Lys Ala Trp Asn Ile Leu Ala 45 Leu Asp Gly Ser Asn Trp Gln Arg Ile Asp Leu Phe Asn Phe Gln Ile 50 60 Asp Val Glu Gly Arg Val Val Glu Asn Ile Ser Lys Arg Cys Gly Gly 65 75 80 Phe Leu Arg Lys Leu Ser Leu Arg Gly Cys Ile Gly Val Gly Asp Ser 85 90 95 Ser Leu Lys Thr Phe Ala Gln Asn Cys Arg Asn Ile Glu His Leu Asn 100 105 110 Leu Asn Gly Cys Thr Lys Ile Thr Asp Ser Thr Cys Tyr Ser Leu Ser 115 120 125 Arg Phe Cys Ser Lys Leu Lys His Leu Asp Leu Thr Ser Cys Val Ser 130 135 140 Ile Thr Asn Ser Ser Leu Lys Gly Ile Ser Glu Gly Cys Arg Asn Leu 145 150 155 160 Glu Tyr Leu Asn Leu Ser Trp Cys Asp Gln Ile Thr Lys Asp Gly Ile 165 170 175 Glu Ala Leu Val Arg Gly Cys Arg Gly Leu Lys Ala Leu Leu Leu Arg 180 185 190 Gly Cys Thr Gln Leu Glu Asp Glu Ala Leu Lys His Ile Gln Asn Tyr 195 200 205 Cys His Glu Leu Val Ser Leu Asn Leu Gln Ser Cys Ser Arg Ile Thr 210 215 220 Asp Glu Gly Val Val Gln Ile Cys Arg Gly Cys His Arg Leu Gln Ala 225 230 235 240 Leu Cys Leu Ser Gly Cys Ser Asn Leu Thr Asp Ala Ser Leu Thr Ala 245 250 255 Leu Gly Leu Asn Cys Pro Arg Leu Gln Ile Leu Glu Ala Arg Cys 260 265 270 Ser His Leu Thr Asp Ala Gly Phe Thr Leu Leu Ala Arg Asn Cys His 275 280 285 Glu Leu Glu Lys Met Asp Leu Glu Glu Cys Ile Leu Ile Thr Asp Ser 290 295 300 Thr Leu Ile Gln Leu Ser Ile His Cys Pro Lys Leu Gln Ala Leu Ser 305 310 315 320 Leu Ser His Cys Glu Leu Ile Thr Asp Asp Gly Ile Leu His Leu Ser 325 330 335

<210> 90 <211> 423 <212> PRT <213> Homo sapiens

 4400> 90 Met 1 val Phe Ser
 Asn Asn Asn Asp Glu Gly Leu Ile Asn Lys Lys Leu Pro 10
 Lys Glu Leu Leu Leu Leu Arg Ile Phe Ser Phe Leu Asp Ile Val Thr Leu 30
 Thr Leu 20

 Lys Arg Cys Arg Cys Ala Gln Ile Ser Lys Arg Cys Arg Sys Ala Gln Trp Gln Arg Ile Asp Gly Ser Asn Trp Gln Arg Ile Asp Leu Phe Asn Phe Gln Thr Asp Val Gly Gly Arg Val Val Glu Asn Ile Ser Lys Arg Cys Gly Gly Phe Leu Robert Cys Lys Leu Ser Leu Arg Gly Cys Ile Gly Val Gly Asp Ser Ser Leu Robert Cys Thr Phe Ala Gln Asn Cys Arg Arg Asn Ile Glu His Leu Asn Leu Asn Leu Asn Cys Robert Cys Ile Gly Val Gly His Leu Asn Ile Cys Trp Ser Leu Asn Ile Cys Ile Gly Cys Thr Lys Ile Thr Asp Ser Thr Cys Trp Ser Leu Ser Arg Phe Ileu Cys Ile Gly Cys Asp Ser Ser Leu Cys Ileu Cys Ileu Gly Cys Arg Asn Leu Glu Trp Ileu Cys I

Glu Leu Val Ser Leu Asn Leu Gln Ser Cys Ser Arg Ile Thr Asp Glu 210 215 220 Gly Val Val Gln Ile Cys Arg Gly Cys His Arg Leu Gln Ala Leu Cys 225 230 235 240 Leu Ser Gly Cys Ser Asn Leu Thr Asp Ala Ser Leu Thr Ala Leu Gly 245 250 255 Leu Asn Cys Pro Arg Leu Gln Ile Leu Glu Ala Ala Arg Cys Ser His 260 265 270 Leu Thr Asp Ala Gly Phe Thr Leu Leu Ala Arg Asn Cys His Glu Leu 275 280 285 Glu Lys Met Asp Leu Glu Glu Cys Ile Leu Ile Thr Asp Ser Thr Leu 290 295 300 Ile Gln Leu Ser Ile His Cys Pro Lys Leu Gln Ala Leu Ser Leu Ser 305 310 315 320 His Cys Glu Leu Ile Thr Asp Asp Gly Ile Leu His Leu Ser Asn Ser 325 330 335 Thr Cys Gly His Glu Arg Leu Arg Val Leu Glu Leu Asp Asn Cys Leu 340 345 350 Leu Ile Thr Asp Val Ala Leu Glu His Leu Glu Asn Cys Arg Gly Leu 355 360 365 Glu Arg Leu Glu Leu Tyr Asp Cys Gln Gln Val Thr Arg Ala Gly Ile 370 375 380 Lys Arg Met Arg Ala Gln Leu Pro His Val Lys Val His Ala Tyr Phe 385 390 395 400 Ala Pro Val Thr Pro Pro Thr Ala Val Ala Gly Ser Gly Gln Arg Leu 405 410 415 Cys Arg Cys Cys Val Ile Leu 420

<210> 91 <211> 46

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: F-box domain sequence

<400> 91

Phe Ser Leu Leu Arg Leu Pro Asp Asp Leu Leu Glu Lys Ile Leu Ser 1 10 15

Arg Leu Pro Leu Lys Asp Leu Leu Ser Leu Ser Lys Val Ser Lys Lys 20 25 30

Phe Arg Ser Leu Val Asp Ser Leu Leu Asp Val Lys Leu Leu 35 40 45

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<210> 92
<211> 172
<212> PRT
<213> Homo sapiens
```

<210> 93 <211> 171 <212> PRT <213> Mus musculus

Arg Gly Val Ala Lys Met Ser Leu Asp Pro Ala Asp Leu Thr His Asp Thr Thr Gly Leu Thr Ala Lys Glu Leu Glu Ala Leu Asp Asp Val Phe Ser Lys Val Tyr Lys Ala Lys Tyr Pro Ile Val Gly Tyr Thr Ala Arg 116 Leu Asp Glu Asp Gly Ser Pro Asn Leu Asp Phe Lys Pro Glu Asp Gln Pro His Phe Asp Ile Lys Asp Glu Phe

<210> 94

<211> 100

<212> PRT

<213> Arabidopsis thaliana

<400> 94

Met Glu Phe Thr Ala Glu Gln Leu Ser Gln Tyr Asn Gly Thr Asp Glu 1 5 10

Ser Lys Pro Ile Tyr Val Ala Ile Lys Gly Arg Val Phe Asp Val Thr 20 25 30

Thr Gly Lys Ser Phe Tyr Gly Ser Gly Gly Asp Tyr Ser Met Phe Ala 35 40 45

Gly Lys Asp Ala Ser Arg Ala Leu Gly Lys Met Ser Lys Asn Glu Glu 50 55 60

Asp Val Ser Pro Ser Leu Glu Gly Leu Thr Glu Lys Glu Ile Asn Thr 65 70 75 80

Leu Asn Asp Trp Glu Thr Lys Phe Glu Ala Lys Tyr Pro Val Val Gly  $85 \hspace{1cm} 90 \hspace{1cm} 95$ 

Arg Val Val Ser 100

<210> 95

<211> 232

<212> PRT

<213> Oryza sativa

<400> 95

Met Ala Ala Ala Val Ala Glu Leu Trp Glu Thr Leu Lys Gln Ala Ile 1 5 10 15

Val Ala Tyr Thr Gly Leu Ser Pro Ala Ala Phe Phe Thr Ala Val Ala 20 25 30

Ala Ala Ala Leu Tyr His Val Val Ser Gly Ile Phe Ala Gly Pro 35 40 45

Pro Pro Pro Pro Pro Arg Pro Arg Asp Glu Pro Glu Ala Glu Pro 50 60 Page 222

Leu Pro Pro Pro Val Gln Leu Gly Glu Val Ser Glu Glu Glu Leu Arg
65 70 75 80 Gln Tyr Asp Gly Ser Asp Pro Lys Lys Pro Leu Leu Met Ala Ile Lys 85 90 95 Gly Gln Ile Tyr Asp Val Thr Gln Ser Arg Met Phe Tyr Gly Pro Gly  $100 \hspace{1cm} 105 \hspace{1cm} 110$ Gly Pro Tyr Ala Leu Phe Ala Gly Lys Asp Ala Ser Arg Ala Leu Ala 115 120 125 Lys Met Ser Phe Glu Pro Gln Asp Leu Thr Gly Asp Ile Ser Gly Leu 130 135 140 Gly Pro Phe Glu Leu Asp Ala Leu Gln Asp Trp Glu Tyr Lys Phe Met 145 150 155 160 Gly Lys Tyr Val Lys Val Gly Thr Val Lys Lys Thr Val Pro Val Glu 165 170 175 Asp Gly Ala Pro Ser Thr Ser Pro Glu Thr Thr Glu Thr Ala Ala Ala 180 185 190 Ala Glu Pro Glu Lys Ala Pro Ala Thr Glu Glu Lys Pro Arg Glu Val 195 200 205 Ser Ser Glu Glu Val Lys Glu Lys Glu Asp Ala Val Ala Ala Ala 210 215 220 Pro Asp Glu Gly Ala Lys Glu Ser 225 230

<210> 96 <211> 104

<212> PRT

<213> Artificial Sequence

<220> <223> Description of Artificial Sequence: Steroid binding domain sequence

<400> 96 Asp Phe Thr Pro Glu Glu Leu Arg Lys Tyr Asp Gly Ser Asp Glu Asp 1 10 15 Lys Pro Ile Tyr Leu Ala Ile Lys Gly Lys Val Tyr Asp Val Thr Arg 20 25 30 Gly Arg Lys Phe Tyr Gly Pro Gly Gly Pro Tyr Ser Leu Phe Ala Gly
35 40 45 Arg Asp Ala Ser Arg Ala Leu Ala Thr Met Ser Phe Asp Glu Glu Asp 50 55 Leu Lys Asp Ser Asp Glu Glu Ile Asp Asp Leu Ser Asp Leu Ser Ala 65 70 75 80 Asp Glu Leu Glu Ala Leu Arg Glu Trp Glu Thr Lys Phe Lys Ala Lys 90 95 Tyr Pro Val Val Gly Arg Leu Ile

<210> 97 <211> 309 <212> PRT <213> Homo sapiens

<400> 97 Met Glu Ala Leu Ala Leu Val Gly Ala Trp Tyr Thr Ala Arg Lys Ser 1 10 15 Ile Thr Val Ile Cys Asp Phe Tyr Ser Leu Ile Arg Leu His Phe Ile 20 25 30 Pro Arg Leu Gly Ser Arg Ala Asp Leu Ile Lys Gln Tyr Gly Arg Trp 40 45Ala Val Ser Gly Ala Thr Asp Gly Ile Gly Lys Ala Tyr Ala Glu 50 55 Glu Leu Ala Ser Arg Gly Leu Asn Ile Ile Leu Ile Ser Arg Asn Glu 65 70 75 80 Glu Lys Leu Gln Val Val Ala Lys Asp Ile Ala Asp Thr Tyr Lys Val 85 90 95 Glu Thr Asp Ile Ile Val Ala Asp Phe Ser Ser Gly Arg Glu Ile Tyr 100 105 110 Leu Pro Ile Arg Glu Ala Leu Lys Asp Lys Asp Val Gly Ile Leu Val 115 120 125 Asn Asn Val Gly Val Phe Tyr Pro Tyr Pro Gln Tyr Phe Thr Gln Leu 130 135 140 Glu Asp Lys Leu Trp Asp Ile Ile Asn Val Asn Ile Ala Ala 150 160 Ser Leu Met Val His Val Val Leu Pro Gly Met Val Glu Arg Lys Lys 165 170 175 Gly Ala Ile Val Thr Ile Ser Ser Gly Ser Cys Cys Lys Pro Thr Pro  $180 \hspace{1cm} 185 \hspace{1cm} 190$ Gln Leu Ala Ala Phe Ser Ala Ser Lys Ala Tyr Leu Asp His Phe Ser 195 200 205 Arg Ala Leu Gln Tyr Glu Tyr Ala Ser Lys Gly Ile Phe Val Gln Ser 210 215 220 Leu Ile Pro Phe Tyr Val Ala Thr Ser Met Thr Ala Pro Ser Asn Phe 225 230 235 240 Leu His Arg Cys Ser Trp Leu Val Pro Ser Pro Lys Val Tyr Ala His 245 250 255 His Ala Val Ser Thr Leu Gly Ile Ser Lys Arg Thr Thr Gly Tyr Trp 260 265 270

Ser His Ser Ile Gln Phe Leu Phe Ala Gln Tyr Met Pro Glu Trp Leu 275 280 285

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Trp Val Trp Gly Ala Asn Ile Leu Asn Arg Ser Leu Arg Lys Glu Ala

Leu Ser Cys Thr Ala 305

<210> 98 <211> 339

<212> PRT

<213> Drosophila melanogaster

Met Gln Pro Val Leu Glu Val Ser Ile Tyr Thr Leu Leu Lys Met Ala 1 5 10 15

Phe Ile Trp Gln Leu Ile Ser Ala Ala Ile Tyr Leu Val Gly Leu Leu 20 25 30

Thr Ile Gly Val Phe Leu Tyr Asp Asn Leu Lys Ser Leu Val Ser Ile 35 40 45

Ile Lys Ala Val Leu Glu Pro Tyr Phe Gln Pro His Leu Pro Arg Thr 50 55 60

Leu Val Asp Lys Phe Gly Gln Trp Ala Val Val Thr Gly Ala Thr Asp 65 70 75 80

Gly Ile Gly Lys Glu Tyr Ala Arg Glu Leu Ala Arg Gln Gly Ile Asn 85 90 95

Leu Val Leu Ile Ser Arg Thr Lys Glu Lys Leu Ile Ala Val Thr Asn 100 105 110

Glu Ile Glu Ser Gln Tyr Lys Val Lys Thr Lys Trp Ile Ala Ala Asp 115 120 125

Phe Ala Lys Gly Arg Glu Val Tyr Asp Gln Ile Glu Lys Glu Leu Ala 130 140

Gly Ile Asp Val Gly Ile Leu Val Asn Asn Val Gly Met Met Tyr Glu 145 150 155 160

His Pro Glu Ser Leu Asp Leu Val Ser Glu Asp Leu Leu Trp Asn Leu 165 170 175

Leu Thr Val Asn Met Gly Ser Val Thr Met Leu Thr Arg Lys Ile Leu 180 185 190

Pro Gln Met Ile Gly Arg Arg Lys Gly Ala Ile Val Asn Leu Gly Ser 195 200 205

Ser Glu Leu Gln Pro Leu Pro Asn Met Thr Val Tyr Ala Ala Ser 210 215 220

Lys Lys Phe Val Thr Tyr Phe Ser Lys Ala Leu Glu Leu Glu Val Ala 225 230 235 240

Glu His Asn Ile His Val Gln Leu Val Met Pro Asn Phe Val Val Thr 245 250 255

Lys Met Asn Ala Tyr Thr Asp Arg Val Met Gln Gly Gly Leu Phe Phe Page 225

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Pro Asn Ala Tyr Thr Phe Ala Arg Ser Ala Val Phe Thr Leu Gly Lys 275

Thr Ser Glu Thr Asn Gly Phe Trp Thr His Gly Ile Gln Tyr Ala Ile Met Lys Leu Ala Pro Leu 310

Phe Lys Arg Leu Arg Ile Glu Ala Leu Glu Gln Lys Gln Lys Lys Leu Clys Leu Thr

<210> 99 <211> 312 <212> PRT

<213> Homo sapiens

 <400> 99
 Ala Leu Sr
 Ala Leu Pro Ala Ala Gly Phe Leu Tyr Trp Val Gly Ala 15
 Ala Tyr Leu Ala Leu Arg Ile Ser Tyr Ser Leu Phe Thr 30
 Ala Leu Arg Val Trp Gly Val Gly Asn Glu Ala Gly Val Gly Asn Glu Ala Gly Val Gly Pro Gly 45
 Ala Leu Arg Val Trp Ala Val Val Thr Gly Ser Thr Asp Gly Ile Gly Lys 50
 Ala Leu Arg Sel Lys Asp Lys Leu Asp Gly His Gly Met Lys Val Val Leu Ile Ro
 Ala Cly Val Gly Ile Gly Lys Ser Thr Asp Gly Ile Gly Lys Ser Tyr Ala Glu Glu Leu Ala Lys His Gly Met Lys Val Val Leu Ile Ro
 Asp Ser Lys Asp Lys Leu Asp Gln Val Ser Ser Glu Ile Lys Glu 95
 Ala Lys His Gly Met Lys Val Val Leu Ile Ro
 Asp Gly Fro Gly 11
 Asp Gly Fro Gly 90
 Asp Ser Ser Glu Ile Lys Glu 19
 Asp Glu Ile Lys Glu Ile Gly 12
 Asp Glu Ile Lys Glu Ile Gly 12
 Asp Glu Ile Lys Glu Ile Gly Ile Gly Ile Gly Ile Gly Ile Gly 12
 Asp Glu Ile Lys Glu Ile Gly Ile G

ValAsp<br/>210PhePheSerGlnCys<br/>215LeuHisCURA-2221.APP<br/>GluArg<br/>220SerLysGlyValPheValGlnSerValLeuProTyrPheValAlaThrLysLeuAlaLysIleArgLysProThrLeuAspLysProSerProGluThrPhe240LysSerAlaLysThrLeuAspLysSerProSerProGluThrPhePhe240LysSerAlaLysThrValGlyLeuGlnSerArgThrAsnGlyTyrLeuIleHisAlaLeuMetGlySerIleIleSerAsnLeuProSerTrpIleTyrLeuLysIleYalMetAsnHisAsnLysSerThrArgAlaHisTyrLeuLysLysThrLysAsnLysSerThrArgAlaHis

<210> 100

<211> 312

<212> PRT

<213> Anas platyrhynchos

 <400> Met Leu Pro
 Ala Ala Gly Leu Leu Trp Trp Val Gly Ala Leu Gly Ala

 Leu Tyr Ala Ala Val Arg Gly Ala Leu Gly Ala Leu Gly Leu Leu Gly Ala Trp Gly Ala Trp Ala Val Val Trp Gly Ala Trp Ala Val Val Trp Gly Ala Trp Ala Leu Ala Lys Arg Gly Met Lys Val Ala Leu Ile Ser Arg Ala Leu Gly Gly Leu Gly Ala Leu Ala Lys Ala Leu Ala Lys Val Ala Leu Ile Ser Arg Ala Leu Gly Gly Ile Gly Lys Ala Trp Ala Leu Gly Ala Leu Ile Ser Arg Ala Leu Gly Val Glu Val Glu Trp Ala Leu Gly Glu Ala Gly Glu Ile Trp Glu Gln Trp Gly Ala Leu Ile Ser Arg Ala Gly Val Glu Val Ile Trp Ala Ala Gly Leu Glu Gly Leu Glu Gly Leu Gly Val Leu Val Ala Ala Gly Leu Glu Gly Leu Gly Val Leu Val Ala Ala Gly Leu Gly Leu Gly Ile Gly Val Leu Val Ala Ala Cly Trp Pro Glu Trp Pro Gly Met Ile Ala Ile Met Ser Val Clys Lys Met Trp Arg Leu Val Leu Pro Gly Met Leu Gly Ala Ala Cly Ile Ala Ala Gly Ile Ala Ile Leu Ala Ala Gly Ile Ser Ser Ala Ala Gly Met Ile Ala Ala Gly Ala Cly Ala

<210> 101 <211> 312

<212> PRT

<213> Mus musculus

ValGluArgSer 180LysGlyValIleLeu 185AsnIleSerSerAla 205SerGlyMetLeuProLeuLeuThrIleTyrSerAlaThrLysAlaPheValAsp 210PhePheSerGlnCys 215LeuHisGluGluTyrLysSerLysGlyIlePheValGlnSerValMetProTyrLeuValAlaThrLysLeuAlaLysIleGlnLysProThrLeuAspLysProSerAlaGluThrPheValLysSerAlaIleLysThrValGlyLeuGlnThrArgThrThrPheValLysSerAlaIleMetGlySerIleAsnSerIleMetPhePheSerIleMetProArgTrpMetTyrPheLysLysArgLysLysAsnSerLysSerLeuArgAsnArgTyrLeuLysLysArgLysAsnSerLysSerLeuArgAsnArg

<210> 102

<211> 271

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Short Chain Alcohol Dehydrogenase (adh\_short) domain sequence

 Adooble 102
 Thr Gly Lys
 Val
 Ala Leu Val
 Thr Gly Ala Ser Ser Gly
 Ile Gly Leu 15
 Ala 16
 Ala 17
 Ala 17
 Ala 17
 Ala 17
 Ala 18
 <t

<210> 103 <211> 1961 <212> PRT <213> Homo sapiens

<400>\_103\_\_\_\_\_

Met Ala Gln Gln Ala Ala Asp Lys Tyr Leu Tyr Val Asp Lys Asn Phe 15 Phe 16 Phe 15 Phe 16 Phe 16 Phe 16 Phe 15 Phe 16 Phe 1

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Met Met Gln Asp Arg Glu Asp Gln Ser Ile Leu Cys Thr Gly Glu Ser 165 170 175 Gly Ala Gly Lys Thr Glu Asn Thr Lys Lys Val Ile Gln Tyr Leu Ala 180 185 190 Tyr Val Ala Ser Ser His Lys Ser Lys Lys Asp Gln Gly Glu Leu Glu 195 200 205 Arg Gln Leu Leu Gln Ala Asn Pro Ile Leu Glu Ala Phe Gly Asn Ala 210 215 220 Lys Thr Val Lys Asn Asp Asn Ser Ser Arg Phe Gly Lys Phe Ile Arg 225 230 235 240 Ile Asn Phe Asp Val Asn Gly Tyr Ile Val Gly Ala Asn Ile Glu Thr 245 250 255 Tyr Leu Leu Glu Lys Ser Arg Ala Ile Arg Gln Ala Lys Glu Glu Arg 260 265 270 Thr Phe His Ile Phe Tyr Tyr Leu Leu Ser Gly Ala Gly Glu His Leu 275 280 285 Thr Asp Leu Leu Glu Pro Tyr Asn Lys Tyr Arg Phe Leu Ser 290 295 300 Asn Gly His Val Thr Ile Pro Gly Gln Gln Asp Lys Asp Met Phe Gln 305 310 315 320 Glu Thr Met Glu Ala Met Arg Ile Met Gly Ile Pro Glu Glu Gln 325 330 335 Met Gly Leu Leu Arg Val Ile Ser Gly Val Leu Gln Leu Gly Asn Ile 340 345 350 Val Phe Lys Lys Glu Arg Asn Thr Asp Gln Ala Ser Met Pro Asp Asn 365 Thr Ala Ala Gln Lys Val Ser His Leu Leu Gly Ile Asn Val Thr Asp 370 375 380 Phe Thr Arg Gly Ile Leu Thr Pro Arg Ile Lys Val Gly Arg Asp Tyr 385 390 395 400 Val Gln Lys Ala Gln Thr Lys Glu Gln Ala Asp Phe Ala Ile Glu Ala 405 410 415 Leu Ala Lys Ala Thr Tyr Glu Arg Met Phe Arg Trp Leu Val Leu Arg 420 425 430 Ile Asn Lys Ala Leu Asp Lys Thr Lys Arg Gln Gly Ala Ser Phe Ile 435 440 445 Glu Gln Leu Cys Ile Asn Tyr Thr Asn Glu Lys Leu Gln Gln Leu Phe 465 470 475 480 Asn His Thr Met Phe Ile Leu Glu Glu Glu Tyr Gln Arg Glu Gly Page 231

Ile Glu Trp Asn Phe Ile Asp Phe Gly Leu Asp Leu Gln Pro Cys Ile 500 505 510 Asp Leu Ile Glu Lys Pro Ala Gly Pro Pro Gly Ile Leu Ala Leu Leu 515 520 525 Glu Glu Cys Trp Phe Pro Lys Ala Thr Asp Lys Ser Phe Val Glu 530 540 Lys Val Met Gln Glu Gln Gly Thr His Pro Lys Phe Gln Lys Pro Lys 545 550 555 560 Gln Leu Lys Asp Lys Ala Asp Phe Cys Ile Ile His Tyr Ala Gly Lys 565 570 575 Val Asp Tyr Lys Ala Asp Glu Trp Leu Met Lys Asn Met Asp Pro Leu 580 585 590 Asn Asp Asn Ile Ala Thr Leu Leu His Gln Ser Ser Asp Lys Phe Val 595 600 605 Ser Glu Leu Trp Lys Asp Val Asp Arg Ile Ile Gly Leu Asp Gln Val 610 620 Ala Gly Met Ser Glu Thr Ala Leu Pro Gly Ala Phe Lys Thr Arg Lys 625 630 635 640 Gly Met Phe Arg Thr Val Gly Gln Leu Tyr Lys Glu Gln Leu Ala Lys 645 650 655 Leu Met Ala Ser Leu Arg Asn Thr Asn Pro Asn Phe Val Arg Cys Ile 660 670 Ile Pro Asn His Glu Lys Lys Ala Gly Lys Leu Asp Pro His Leu Val 675 680 685 Leu Asp Gln Leu Arg Cys Asn Gly Val Leu Glu Gly Ile Arg Ile Cys 690 695 700 Arg Gln Gly Phe Pro Asn Arg Val Val Phe Gln Glu Phe Arg Gln Arg 705 710 715 720 Tyr Glu Ile Leu Thr Pro Asn Ser Ile Pro Lys Gly Phe Met Asp Gly 725 730 735 Lys Gln Ala Cys Val Leu Met Ile Lys Ala Leu Glu Leu Asp Ser Asn 740 745 750 Leu Tyr Arg Ile Gly Gln Ser Lys Val Phe Phe Arg Ala Gly Val Leu 755 760 765 Ala His Leu Glu Glu Glu Arg Asp Leu Lys Ile Thr Asp Val Ile Ile 770 780 Gly Phe Gln Ala Cys Cys Arg Gly Tyr Leu Ala Arg Lys Ala Phe Ala 785 790 795 800 Lys Arg Gln Gln Leu Thr Ala Met Lys Val Leu Gln Arg Asn Cys 805 810 815 Ala Ala Tyr Leu Lys Leu Arg Asn Trp Gln Trp Trp Arg Leu Phe Thr Page 232

830

Lys Val Lys Pro Leu Leu Gln Val Ser Arg Gln Glu Glu Met Met 835 840 845 840 Ala Lys Glu Glu Leu Val Lys Val Arg Glu Lys Gln Leu Ala Ala 850 855 860 Glu Asn Arg Leu Met Glu Met Glu Thr Leu Gln Ser Gln Leu Met Ala 865 870 875 880 Glu Lys Leu Gln Leu Gln Glu Gln Leu Gln Ala Glu Thr Glu Leu Cys 890 Ala Glu Ala Glu Glu Leu Arg Ala Arg Leu Thr Ala Lys Lys Gln Glu 900 905 910 Leu Glu Glu Ile Cys His Asp Leu Glu Ala Arg Val Glu Glu Glu 915 920 925 Glu Arg Tyr Gln His Leu Gln Ala Glu Lys Lys Lys Met Gln Gln Asn 930 935 940 Ile Gln Glu Leu Glu Glu Gln Leu Glu Glu Glu Ser Ala Arg Gln 945 950 955 960 Lys Leu Gln Leu Glu Lys Val Thr Thr Glu Ala Lys Leu Lys Lys Leu 965 970 975 Glu Glu Glu Gln Ile Ile Leu Glu Asp Gln Asn Cys Lys Leu Ala Lys 980 985 990 Glu Lys Lys Leu Leu Glu Asp Arg Ile Ala Glu Phe Thr Thr Asn Leu 995 1000 1005 Thr Glu Glu Glu Lys Ser Lys Ser Leu Ala Lys Leu Lys Asn Lys 1010 1015 1020 His Glu Ala Met Ile Thr Asp Leu Glu Glu Arg Leu Arg Arg Glu Glu 1025 1030 1035 1040 1025 1030 Lys Gln Arg Gln Glu Leu Glu Lys Thr Arg Arg Lys Leu Glu Gly Asp 1045 1050 1055 Ser Thr Asp Leu Ser Asp Gln Ile Ala Glu Leu Gln Ala Gln Ile Ala 1060 1065 1070 1060 Glu Leu Lys Met Gln Leu Ala Lys Lys Glu Glu Glu Leu Gln Ala Ala 1075 1080 1085 Leu Ala Arg Val Glu Glu Glu Ala Ala Gln Lys Asn Met Ala Leu Lys 1095 1100 Lys Ile Arg Glu Leu Glu Ser Gln Ile Ser Glu Leu Gln Glu Asp Leu 1105 1110 1115 1120 Glu Ser Glu Arg Ala Ser Arg Asn Lys Ala Glu Lys Gln Lys Arg Asp 1125 1130 1135 Leu Gly Glu Glu Leu Glu Ala Leu Lys Thr Glu Leu Glu Asp Thr Leu 1140 1145 1150 Asp Ser Thr Ala Ala Gln Gln Glu Leu Arg Ser Lys Arg Glu Gln Glu Page 233

1160

Val Asn Ile Leu Lys Lys Thr Leu Glu Glu Glu Ala Lys Thr His Glu 1170 1175 1180

Leu Ala Glu Gln Leu Glu Gln Thr Lys Arg Val Lys Ala Asn Leu Glu 1205 1210 1215

Lys Ala Lys Gln Thr Leu Glu Asn Glu Arg Gly Glu Leu Ala Asn Glu 1220 1225 1230

Val Lys Val Leu Leu Gln Gly Gly Arg Asp Ser Glu His Lys Arg Lys 1235 1240 1245

Lys Val Glu Ala Gln Leu Gln Glu Leu Gln Val Lys Phe Asn Glu Gly 1250 1255 1260

Glu Arg Val Arg Thr Glu Leu Ala Asp Lys Val Thr Lys Leu Gln Val 1265 1270 1275 1280

Glu Leu Asp Asn Val Thr Gly Leu Leu Ser Gln Ser Asp Ser Lys Ser 1285 1290 1295

Ser Lys Leu Thr Lys Asp Phe Ser Ala Leu Glu Ser Gln Leu Gln Asp 1300 1305 1310

Thr Gln Glu Leu Leu Gln Glu Glu Asn Arg Gln Lys Leu Ser Leu Ser 1315 1320 1325

Thr Lys Leu Lys Gln Val Glu Asp Glu Lys Asn Ser Phe Arg Glu Gln 1330 1335 1340

Leu Glu Glu Glu Glu Glu Ala Lys His Asn Leu Glu Lys Gln Ile 1345 1350 1355 1360

Ala Thr Leu His Ala Gln Val Ala Asp Met Lys Lys Lys Met Glu Asp 1365 1370 1375

Ser Val Gly Cys Leu Glu Thr Ala Glu Glu Val Lys Arg Lys Leu Gln 1380 1385 1390

Lys Asp Leu Glu Gly Leu Ser Gln Arg His Glu Glu Lys Val Ala Ala 1395 1400 1405

Tyr Asp Lys Leu Glu Lys Thr Lys Thr Arg Leu Gln Glu Leu Asp 1410 1415 1420

Asp Leu Leu Val Asp Leu Asp His Gln Arg Gln Ser Ala Cys Asn Leu 1425 1430 1435 1440

Glu Lys Lys Gln Lys Lys Phe Asp Gln Leu Leu Ala Glu Glu Lys Thr 1445 1450 1455

Ile Ser Ala Lys Tyr Ala Glu Glu Arg Asp Arg Ala Glu Ala 1460 1465 1470

Arg Glu Lys Glu Thr Lys Ala Leu Ser Leu Ala Arg Ala Leu Glu Glu 1475 1480 1485

Ala Met Glu Gln Lys Ala Glu Leu Glu Arg Leu Asn Lys Gln Phe Arg Page 234

Thr Glu Met Glu Asp Leu Met Ser Ser Lys Asp Asp Val Gly Lys Ser 1505 1510 1520

Val His Glu Leu Glu Lys Ser Lys Arg Ala Leu Glu Gln Gln Val Glu 1525 1530 1535

Glu Met Lys Thr Gln Leu Glu Glu Leu Glu Asp Glu Leu Gln Ala Thr 1540 1545 1550

Glu Asp Ala Lys Leu Arg Leu Glu Val Asn Leu Gln Ala Met Lys Ala 1555 1560 1565

Gln Phe Glu Arg Asp Leu Gln Gly Arg Asp Glu Gln Ser Glu Glu Lys 1570 1580

Lys Lys Gln Leu Val Arg Gln Val Arg Glu Met Glu Ala Glu Leu Glu 1585 1590 1595 1600

Asp Glu Arg Lys Gln Arg Ser Met Ala Val Ala Ala Arg Lys Leu 1605 1610 1615

Glu Met Asp Leu Lys Asp Leu Glu Ala His Ile Asp Ser Ala Asn Lys 1620 1625 1630

Asn Arg Asp Glu Ala Ile Lys Gln Leu Arg Lys Leu Gln Ala Gln Met 1635 1640 1645

Lys Asp Cys Met Arg Glu Leu Asp Asp Thr Arg Ala Ser Arg Glu Glu 1650 1655 1660

Ile Leu Ala Gln Ala Lys Glu Asn Glu Lys Lys Leu Lys Ser Met Glu 1665 1670 1675 1680

Ala Glu Met Ile Gln Leu Gln Glu Glu Leu Ala Ala Glu Arg Ala 1685 1690 1695

Lys Arg Gln Ala Gln Gln Glu Arg Asp Glu Leu Ala Asp Glu Ile Ala 1700 1705 1710

Asn Ser Ser Gly Lys Gly Ala Leu Ala Leu Glu Glu Lys Arg Arg Leu 1715 1720 1725

Glu Ala Arg Ile Ala Gln Leu Glu Glu Glu Glu Glu Glu Gln Gly 1730 1735 1740

Asn Thr Glu Leu Ile Asn Asp Arg Leu Lys Lys Ala Asn Leu Gln Ile 1745 1750 1760

Asp Gln Ile Asn Ala Asp Leu Asn Leu Glu Arg Gly His Ala Gln Lys 1765 1770 1775

Asn Glu Asn Ala Arg Gln Gln Leu Glu Arg Gln Asn Lys Glu Leu Lys 1780 1785 1790

Val Lys Leu Gln Glu Met Glu Gly Thr Val Lys Ser Lys Tyr Lys Ala 1795 1800 1805

Ser Ile Thr Ala Leu Glu Ala Lys Ile Ala Gln Leu Glu Glu Gln Leu 1810 1815 1820

Asp Asn Glu Thr Lys Glu Arg Gln Ala Ala Cys Lys Gln Val Arg Arg Page 235 Thr Glu Lys Lys Leu Lys Asp Val Leu Leu Gln Val Asp Asp Glu Arg 1845 1850 1855

Arg Asn Ala Glu Gln Tyr Lys Asp Gln Ala Asp Lys Ala Ser Thr Arg 1860 1865 1870

Leu Lys Gln Leu Lys Arg Gln Leu Glu Glu Ala Glu Glu Glu Ala Gln 1875 1880 1885

Arg Ala Asn Ala Ser Arg Arg Lys Leu Gln Arg Glu Leu Glu Asp Ala 1890 1895 1900

Thr Glu Thr Ala Asp Ala Met Asn Arg Glu Val Ser Ser Leu Lys Asn 1905 1910 1915 1920

Lys Leu Arg Arg Gly Asp Leu Pro Phe Val Val Pro Arg Arg Met Ala 1925 1930 1935

Arg Lys Gly Ala Gly Asp Gly Ser Asp Glu Glu Val Asp Gly Lys Ala 1940 1945 1950

Asp Gly Ala Glu Ala Lys Pro Ala Glu 1955 1960

<210> 104

<211> 1960

<212> PRT

<213> Homo sapiens

<400> 104

Met Ala Gln Gln Ala Ala Asp Lys Tyr Leu Tyr Val Asp Lys Asn Phe

1 10 15

Ile Asn Asn Pro Leu Ala Gln Ala Asp Trp Ala Ala Lys Lys Leu Val $20 \hspace{1cm} 25 \hspace{1cm} 30$ 

Trp Val Pro Ser Asp Lys Ser Gly Phe Glu Pro Ala Ser Leu Lys Glu
35 40 45

Glu Val Gly Glu Glu Ala Ile Val Glu Leu Val Glu Asn Gly Lys Lys 50 55 60

Val Lys Val Asn Lys Asp Asp Ile Gln Lys Met Asn Pro Pro Lys Phe 65 70 75 80

Ser Lys Val Glu Asp Met Ala Glu Leu Thr Cys Leu Asn Glu Ala Ser 85 90 95

Val Leu His Asn Leu Lys Glu Arg Tyr Tyr Ser Gly Leu Ile Tyr Thr  $100 \hspace{1cm} 105 \hspace{1cm} 110$ 

Tyr Ser Gly Leu Phe Cys Val Val Ile Asn Pro Tyr Lys Asn Leu Pro 115 120 125

Ile Tyr Ser Glu Glu Ile Val Glu Met Tyr Lys Gly Lys Lys Arg His 130 135 140

Glu Met Pro Pro His Ile Tyr Ala Ile Thr Asp Thr Ala Tyr Arg Ser 145 150 155 160 CURA2221.APP
Met Met Gln Asp Arg Glu Asp Gln Ser Ile Leu Cys Thr Gly Glu Ser
165 170 175 Gly Ala Gly Lys Thr Glu Asn Thr Lys Lys Val Ile Gln Tyr Leu Ala 180 185 190 Tyr Val Ala Ser Ser His Lys Ser Lys Lys Asp Gln Gly Glu Leu Glu 195 200 205 Arg Gln Leu Leu Gln Ala Asn Pro Ile Leu Glu Ala Phe Gly Asn Ala 210 215 220 Lys Thr Val Lys Asn Asp Asn Ser Ser Arg Phe Gly Lys Phe Ile Arg 225 230 235 240 Ile Asn Phe Asp Val Asn Gly Tyr Ile Val Gly Ala Asn Ile Glu Thr 245 250 255 Tyr Leu Leu Glu Lys Ser Arg Ala Ile Arg Gln Ala Lys Glu Glu Arg 260 265 270 Thr Phe His Ile Phe Tyr Tyr Leu Leu Ser Gly Ala Gly Glu His Leu 275 280 285 Lys Thr Asp Leu Leu Glu Pro Tyr Asn Lys Tyr Arg Phe Leu Ser 290 295 300 Asn Gly His Val Thr Ile Pro Gly Gln Gln Asp Lys Asp Met Phe Gln 305 310 315 320 Glu Thr Met Glu Ala Met Arg Ile Met Gly Ile Pro Glu Glu Gln 325 330 335 Met Gly Leu Leu Arg Val Ile Ser Gly Val Leu Gln Leu Gly Asn Ile 340 350 Val Phe Lys Lys Glu Arg Asn Thr Asp Gln Ala Ser Met Pro Asp Asn 355 360 365 Thr Ala Ala Gln Lys Val Ser His Leu Leu Gly Ile Asn Val Thr Asp 370 380 Phe Thr Arg Gly Ile Leu Thr Pro Arg Ile Lys Val Gly Arg Asp Tyr 385 390 395 400 Val Gln Lys Ala Gln Thr Lys Glu Gln Ala Asp Phe Ala Ile Glu Ala 405 410 415 Leu Ala Lys Ala Thr Tyr Glu Arg Met Phe Arg Trp Leu Val Leu Arg 420 425 430 Ile Asn Lys Ala Leu Asp Lys Thr Lys Arg Gln Gly Ala Ser Phe Ile 435 440 445Glu Gln Leu Cys Ile Asn Tyr Thr Asn Glu Lys Leu Gln Gln Leu Phe 465 470 475 480 Asn His Thr Met Phe Ile Leu Glu Glu Glu Glu Tyr Gln Arg Glu Gly 485 490 495 CURA2221.APP

Ile Glu Trp Asn Phe Ile Asp Phe Gly Leu Asp Leu Gln Pro Cys Ile
500 505 Asp Leu Ile Glu Lys Pro Ala Gly Pro Pro Gly Ile Leu Ala Leu Leu 515 520 525 Asp Glu Glu Cys Trp Phe Pro Lys Ala Thr Asp Lys Ser Phe Val Glu 530 535 540 Lys Val Met Gln Glu Gln Gly Thr His Pro Lys Phe Gln Lys Pro Lys 545 550 560 Gln Leu Lys Asp Lys Ala Asp Phe Cys Ile Ile His Tyr Ala Gly Lys 565 570 575 Val Asp Tyr Lys Ala Asp Glu Trp Leu Met Lys Asn Met Asp Pro Leu
580 590 Asn Asp Asn Ile Ala Thr Leu Leu His Gln Ser Ser Asp Lys Phe Val 595 600 605 Ser Glu Leu Trp Lys Asp Val Asp Arg Ile Ile Gly Leu Asp Gln Val 610 615 620 Ala Gly Met Ser Glu Thr Ala Leu Pro Gly Ala Phe Lys Thr Arg Lys 625 630 635 640 Gly Met Phe Arg Thr Val Gly Gln Leu Tyr Lys Glu Gln Leu Ala Lys 645 650 655 Leu Met Ala Thr Leu Arg Asn Thr Asn Pro Asn Phe Val Arg Cys Ile 660 665 670 Ile Pro Asn His Glu Lys Lys Ala Gly Lys Leu Asp Pro His Leu Val 675 680 685 Leu Asp Gln Leu Arg Cys Asn Gly Val Leu Glu Gly Ile Arg Ile Cys 690 695 700 Arg Gln Gly Phe Pro Asn Arg Val Val Phe Gln Glu Phe Arg Gln Arg 705 710 715 720 Tyr Glu Ile Leu Thr Pro Asn Ser Ile Pro Lys Gly Phe Met Asp Gly 725 730 735 Lys Gln Ala Cys Val Leu Met Ile Lys Ala Leu Glu Leu Asp Ser Asn 740 745 750 Leu Tyr Arg Ile Gly Gln Ser Lys Val Phe Phe Arg Ala Gly Val Leu 755 760 765 Ala His Leu Glu Glu Glu Arg Asp Leu Lys Ile Thr Asp Val Ile Ile 770 780 Gly Phe Gln Ala Cys Cys Arg Gly Tyr Leu Ala Arg Lys Ala Phe Ala 785 790 795 800 Lys Arg Gln Gln Leu Thr Ala Met Lys Val Leu Gln Arg Asn Cys 805 810 815 Ala Ala Tyr Leu Lys Leu Arg Asn Trp Gln Trp Trp Arg Leu Phe Thr 820 825 830

CURA2221.APP Lys Val Lys Pro Leu Leu Gln Val Ser Arg Gln Glu Glu Met Met 835 840 845 840 Ala Lys Glu Glu Leu Val Lys Val Arg Glu Lys Gln Leu Ala Ala 850 855 860 Glu Asn Arg Leu Thr Glu Met Glu Thr Leu Gln Ser Gln Leu Met Ala 865 870 875 880 Glu Lys Leu Gln Leu Gln Glu Gln Leu Gln Ala Glu Thr Glu Leu Cys Ala Glu Ala Glu Glu Leu Arg Ala Arg Leu Thr Ala Lys Lys Gln Glu 900 905 910 Leu Glu Glu Ile Cys His Asp Leu Glu Ala Arg Val Glu Glu Glu 915 920 925 Glu Arg Cys Gln His Leu Gln Ala Glu Lys Lys Lys Met Gln Gln Asn 930 935 940 Ile Gln Glu Leu Glu Glu Gln Leu Glu Glu Glu Glu Ser Ala Arg Gln 945 950 955 960 Lys Leu Gln Leu Glu Lys Val Thr Thr Glu Ala Lys Leu Lys Lys Leu 965 970 975 Glu Glu Gln Ile Ile Leu Glu Asp Gln Asn Cys Lys Leu Ala Lys 980 985 990 Glu Lys Lys Leu Leu Glu Asp Arg Ile Ala Glu Phe Thr Thr Asn Leu 995 1005 Thr Glu Glu Glu Lys Ser Lys Ser Leu Ala Lys Leu Lys Asn Lys 1010 1015 1020 His Glu Ala Met Ile Thr Asp Leu Glu Glu Arg Leu Arg Arg Glu Glu 1025 1030 1035 1040 1030 Lys Gln Arg Gln Glu Leu Glu Lys Thr Arg Arg Lys Leu Glu Gly Asp  $1045 \hspace{1cm} 1050 \hspace{1cm} 1055$ Ser Thr Asp Leu Ser Asp Gln Ile Ala Glu Leu Gln Ala Gln Ile Ala 1060 1065 1070 Glu Leu Lys Met Gln Leu Ala Lys Lys Glu Glu Glu Leu Gln Ala Ala 1075 1080 1085 Leu Ala Arg Val Glu Glu Glu Ala Ala Gln Lys Asn Met Ala Leu Lys 1100 Lys Ile Arg Glu Leu Glu Ser Gln Ile Ser Glu Leu Gln Glu Asp Leu 1115 1105 1110 Glu Ser Glu Arg Ala Ser Arg Asn Lys Ala Glu Lys Gln Lys Arg Asp 1125 1130 1135 Leu Gly Glu Glu Leu Glu Ala Leu Lys Thr Glu Leu Glu Asp Thr Leu 1140 1145 1150 Asp Ser Thr Ala Ala Gln Gln Glu Leu Arg Ser Lys Arg Glu Gln Glu

val	Asn l170	Ile	Leu	Lys		Thr 1175	Leu	Glu		Glu	1.API Ala 1180		Thr	His	Glu
Ala 1185		Ile	Gln		Met 1190	Arg	Gln	Lys		Ser 1195	Gln	Ala	val	Glu E	G]u 1200
Leu	Ala	Glu		Leu 1205	Glu	Gln	Thr		Arg 1210	۷al	Lys	Аlа		Leu 1215	Glu
Lys	Ala		Gln 1220	Thr	Leu	Glu		G]u 1225	Arg	Glу	Glu		А]а 1230	Asn	Glu
۷al		Va1 1235	Leu	Leu	Gln	Gly :	Lys 1240	Gly	Asp	Ser		His L245	Lys	Arg	Lys
	Va 1 L250	Glu	Ala	Gln		G]n 1255	Glu	Leu	Gln		Lys 1260	Phe	Asn	Glu	Gly
Glu 1269	Arg	۷al	Arg		G1u 1270	Leu	Ala	Asp	Lys	Va1 1275	Thr	Lys	Leu	G]n	va1 1280
Glu	Leu	Asp		val 1285	Thr	Gly	Leu		Ser 1290	Gln	Ser	Asp		Lys 1295	Ser
Ser	Lys		Thr 1300		Asp	Phe		Ala 1305	Leu	Glu	Ser		Leu 1310	Gln	Asp
Thr		G]u 1315	Leu	Leu	Gln		G]u 1320	Asn	Arg	G∏n		Leu 1325	Ser	Leu	Ser
	Lys L330	Leu	Lys	Gln		G] u 1335	Asp	Glu	Lys		Ser 1340	Phe	Arg	Glu	G∏n
Leu 1345		Glu	Glu		Glu 1350	Ala	Lys	His		Leu 1355	Glu	Lys	Gln	Ile	а]а 1360
Thr	Leu	His		G]n 1365	val	Ala	Asp		Lys 1370	Lys	Lys	Met		Asp 1375	Ser
Val	Gly		Leu 1380	Glu	Thr	Ala		G] u 1385	val	Lys	Arg		Leu 1390	Gln	Lys
Asp		G]u 1395	Gly	Leu	Ser		Arg 1400	His	Glu	Glu		Va1 L405	Ala	Аlа	Tyr
Asp	Lys 1410	Leu	Glu	Lys	Thr	Lys 1415	Thr	Arg	Leu	Gln	G]n 1420	Glu	Leu	Asp	Asp
Leu 142		۷al	Asp		Asp 1430	His	Gln	Arg		Ser 1435	Ala	Cys	Asn	Leu 1	Glu L440
Lys	Lys	Gln		Lys 1445	Phe	Asp	Gln		Leu 1450	Ala	Glu	Glu		Thr 1455	Ile
Ser	Ala		туг 1460		Glu	Glu		Asp 1465	Arg	Аlа	Glu		Glu 1470	Ala	Arg
Glu		G]u 1475	Thr	Lys	Аlа	Leu	Ser 1480	Leu	Ala	Arg	Ala	Leu L485	Glu	Glu	Ala

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His Glu Leu Glu Lys Ser Lys Arg Ala Leu Glu Gln Gln Val Glu Glu 1525 1530 1535

Met Lys Thr Gln Leu Glu Glu Leu Glu Asp Glu Leu Gln Ala Thr Glu 1540 1545 1550

Asp Ala Lys Leu Arg Leu Glu Val Asn Leu Gln Ala Met Lys Ala Gln 1555 1560 1565

Phe Glu Arg Asp Leu Gln Gly Arg Asp Glu Gln Ser Glu Glu Lys Lys 1570 1575 1580

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Arg Asp Glu Ala Ile Lys Gln Leu Arg Lys Leu Gln Ala Gln Met Lys 1635 1640 1645

Asp Cys Met Arg Glu Leu Asp Asp Thr Arg Ala Ser Arg Glu Glu Ile 1650 1655 1660

Leu Ala Gln Ala Lys Glu Asn Glu Lys Lys Leu Lys Ser Met Glu Ala 1665 1670 1675 1680

Glu Met Ile Gln Leu Gln Glu Glu Leu Ala Ala Glu Arg Ala Lys 1685 1690 1695

Arg Gln Ala Gln Glu Arg Asp Glu Leu Ala Asp Glu Ile Ala Asn 1700 1705 1710

Ser Ser Gly Lys Gly Ala Leu Ala Leu Glu Glu Lys Arg Arg Leu Glu 1715 1720 1725

Ala Arg Ile Ala Gln Leu Glu Glu Glu Leu Glu Glu Glu Gln Gly Asn 1730 1735 1740

Thr Glu Leu Ile Asn Asp Arg Leu Lys Lys Ala Asn Leu Gln Ile Asp 1745 1750 1760

Gln Ile Asn Thr Asp Leu Asn Leu Glu Arg Ser His Ala Gln Lys Asn 1765 1770 1775

Glu Asn Ala Arg Gln Gln Leu Glu Arg Gln Asn Lys Glu Leu Lys Val 1780 1785 1790

Lys Leu Gln Glu Met Glu Gly Thr Val Lys Ser Lys Tyr Lys Ala Ser 1795 1800 1805

Ile Thr Ala Leu Glu Ala Lys Ile Ala Gln Leu Glu Glu Gln Leu Asp 1810 1815 1820

Asn Glu Thr Lys Glu Arg Gln Ala Ala Cys Lys Gln Val Arg Arg Thr 1825 1830 1835 1840

CURA2221.APP Glu Lys Lys Leu Lys Asp Val Leu Leu Gln Val Asp Asp Glu Arg Arg 1845 1850 1855 Asn Ala Glu Gln Tyr Lys Asp Gln Ala Asp Lys Ala Ser Thr Arg Leu 1860 1865 1870 Lys Gln Leu Lys Arg Gln Leu Glu Glu Ala Glu Glu Ala Gln Arg 1875 1880 1885 Ala Asn Ala Ser Arg Arg Lys Leu Gln Arg Glu Leu Glu Asp Ala Thr 1890 1895 1900

Glu Thr Ala Asp Ala Met Asn Arg Glu Val Ser Ser Leu Lys Asn Lys 1905 1910 1915 1920

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35 40 45 Glu Val Gly Glu Glu Ala Ile Val Glu Leu Val Glu Asn Gly Lys Lys 50 55 60 Val Lys Val Asn Lys Asp Asp Ile Gln Lys Met Asn Pro Pro Lys Phe 65 70 75 80 Ser Lys Val Glu Asp Met Ala Glu Leu Thr Cys Leu Asn Glu Ala Ser 85 90 95 Val Leu His Asn Leu Lys Glu Arg Tyr Tyr Ser Gly Leu Ile Tyr Thr 100 105 110Tyr Ser Gly Leu Phe Cys Val Val Ile Asn Pro Tyr Lys Asn Leu Pro 115 120 125 Ile Tyr Ser Glu Glu Ile Val Asp Met Tyr Lys Gly Lys Lys Arg His 130 135 140 Glu Met Pro Pro His Ile Tyr Ala Ile Thr Asp Thr Ala Tyr Arg Ser 145 150 155 160 Met Met Gln Asp Arg Glu Asp Gln Ser Ile Leu Cys Thr Gly Glu Ser 165 170 175

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Gly Ala Gly Lys Thr Glu Asn Thr Lys Lys Val Ile Gln Tyr Leu Ala 180 185 190 His Val Ala Ser Ser His Lys Ser Lys Lys Asp Gln Gly Glu Leu Glu 195 200 205 Arg Gln Leu Cln Ala Asn Pro Ile Leu Glu Ala Phe Gly Asn Ala 210 220 Lys Thr Val Lys Asn Asp Asn Ser Ser Arg Phe Gly Lys Phe Ile Arg 225 230 235 240 Ile Asn Phe Asp Val Asn Gly Tyr Ile Val Gly Ala Asn Ile Glu Thr 245 250 255 Tyr Leu Leu Glu Lys Ser Arg Ala Ile Arg Gln Ala Lys Glu Glu Arg 260 265 270 Thr Phe His Ile Phe Tyr Tyr Leu Leu Ser Gly Ala Gly Glu His Leu 275 280 285 Lys Thr Asp Leu Leu Glu Pro Tyr Asn Lys Tyr Arg Phe Leu Ser 290 295 300 Asn Gly His Val Thr Ile Pro Gly Gln Gln Asp Lys Asp Met Phe Gln 305 310 315 320 Glu Thr Met Glu Ala Met Arg Ile Met Gly Ile Pro Glu Asp Glu Gln 325 330 335 Met Gly Leu Leu Arg Val Ile Ser Gly Val Leu Gln Leu Gly Asn Ile 340 345 350 Val Phe Lys Lys Glu Arg Asn Thr Asp Gln Ala Ser Met Pro Asp Asn 365 Thr Ala Ala Gln Lys Val Ser His Leu Leu Gly Ile Asn Val Thr Asp 370 375 380 Phe Thr Arg Gly Ile Leu Thr Pro Arg Ile Lys Val Gly Arg Asp Tyr 385 390 395 400 Val Gln Lys Ala Gln Thr Lys Glu Gln Ala Asp Phe Ala Ile Glu Ala 405 410 415 Leu Ala Lys Ala Thr Tyr Glu Arg Met Phe Arg Trp Leu Val Leu Arg 420 425 430 Ile Asn Lys Ala Leu Asp Lys Thr Lys Arg Gln Gly Ala Ser Phe Ile 435 440 445 Gly Ile Leu Asp Ile Ala Gly Phe Glu Ile Phe Asp Leu Asn Ser Phe 450 455 460 Glu Gln Leu Cys Ile Asn Tyr Thr Asn Glu Lys Leu Gln Gln Leu Phe 465 470 475 480 Asn His Thr Met Phe Ile Leu Glu Gln Glu Glu Tyr Gln Arg Glu Gly 485 490 495 Ile Glu Trp Asn Phe Ile Asp Phe Gly Leu Asp Leu Gln Pro Cys Ile 500 505 510Page 243

Asp Leu Ile Glu Lys Pro Ala Gly Pro Pro Gly Ile Leu Ala Leu Leu 515 520 525 Asp Glu Glu Cys Trp Phe Pro Lys Ala Thr Asp Lys Ser Phe Val Glu 530 540 Lys Val Val Gln Glu Gln Gly Thr His Pro Lys Phe Gln Lys Pro Lys 545 550 560 Gln Leu Lys Asp Lys Ala Asp Phe Cys Ile Ile His Tyr Ala Gly Lys 565 570 575 Val Asp Tyr Lys Ala Asp Glu Trp Leu Met Lys Asn Met Asp Pro Leu 580 585 590 Asn Asp Asn Ile Ala Thr Leu Leu His Gln Ser Ser Asp Lys Phe Val 595 600 605 Ser Glu Leu Trp Lys Asp Val Asp Arg Ile Ile Gly Leu Asp Gln Val 610 620 Ala Gly Met Ser Glu Thr Ala Leu Pro Gly Ala Phe Lys Thr Arg Lys 625 630 635 640 Gly Met Phe Arg Thr Val Gly Gln Leu Tyr Lys Glu Gln Leu Ala Lys 645 650 655 Leu Met Ala Thr Leu Arg Asn Thr Asn Pro Asn Phe Val Cys Cys Ile 660 665 670 Ile Pro Asn His Glu Lys Lys Ala Gly Lys Leu Asp Pro His Leu Val 675 680 685 Leu Asp Gln Leu Arg Cys Asn Gly Val Leu Glu Gly Ile Arg Ile Cys 690 695 700 Arg Gln Gly Phe Pro Asn Arg Val Val Phe Gln Glu Phe Arg Gln Arg 705 710 715 720 Tyr Glu Ile Leu Thr Pro Asn Ser Ile Pro Lys Gly Phe Met Asp Gly  $725 \hspace{1cm} 730 \hspace{1cm} 735$ Lys Gln Ala Cys Val Leu Met Ile Lys Ala Leu Glu Leu Asp Ser Asn 740 745 750 Leu Tyr Arg Ile Gly Gln Ser Lys Val Phe Phe Arg Ser Gly Val Leu 755 760 765 Ala His Leu Glu Glu Glu Arg Asp Leu Lys Ile Thr Asp Val Ile Ile 770 775 780 Gly Phe Gln Ala Cys Cys Arg Gly Tyr Leu Ala Arg Lys Ala Phe Ala 785 790 795 800 Lys Arg Gln Gln Gln Leu Thr Ala Met Lys Val Leu Gln Arg Asn Cys 805 810 815 Ala Ala Tyr Leu Arg Leu Arg Asn Trp Gln Trp Trp Arg Leu Phe Thr 820 825 830 Lys Val Lys Pro Leu Leu Asn Ser Ile Arg His Glu Asp Glu Leu Leu 835 840 845 Page 244

Ala Lys Glu Ala Glu Leu Thr Lys Val Arg Glu Lys His Leu Ala Ala 850 855 860 Glu Asn Arg Leu Thr Glu Met Glu Thr Met Gln Ser Gln Leu Met Ala 865 870 875 880 Glu Lys Leu Gln Leu Gln Glu Gln Leu Gln Ala Lys Thr Glu Leu Cys Ala Glu Ala Glu Glu Leu Arg Ala Arg Leu Thr Ala Lys Lys Gln Glu 900 905 910 Leu Glu Glu Ile Cys His Asp Leu Glu Ala Arg Val Glu Glu Glu 915 920 925 Glu Arg Cys Gln Tyr Leu Gln Ala Glu Lys Lys Lys Met Gln Gln Asn 930 935 940 Ile Gln Glu Leu Glu Glu Glu Glu Glu Glu Ser Ala Arg Gln 945 950 955 960 Lys Leu Gln Leu Glu Lys Val Thr Thr Glu Ala Lys Leu Lys Lys Leu 965 970 975 Glu Glu Asp Gln Ile Ile Met Glu Asp Gln Asn Cys Lys Leu Ala Lys 980 985 990 Glu Lys Lys Leu Glu Asp Arg Val Ala Glu Phe Thr Thr Asp Leu 995 1000 1005 Met Glu Glu Glu Lys Ser Lys Ser Leu Ala Lys Leu Lys Asn Lys 1010 1015 1020 His Glu Ala Met Ile Thr Asp Leu Glu Glu Arg Leu Arg Arg Glu Glu 1025 1030 1035 1040 1030 Lys Gln Arg Gln Glu Leu Glu Lys Thr Arg Arg Lys Leu Glu Gly Asp 1045 1050 1055 Ser Thr Asp Leu Ser Asp Gln Ile Ala Glu Leu Gln Ala Gln Ile Ala 1065 1060 Glu Leu Lys Met Gln Leu Ala Lys Lys Glu Glu Glu Leu Gln Ala Ala 1075 1080 1085 Leu Ala Arg Val Glu Glu Glu Ala Ala Gln Lys Asn Met Ala Leu Lys 1095 Lys Ile Arg Glu Leu Glu Thr Gln Ile Ser Glu Leu Gln Glu Asp Leu 1105 1115 1110 Glu Ser Glu Arg Ala Cys Arg Asn Lys Ala Glu Lys Gln Lys Arg Asp 1125 1130 1135 Leu Gly Glu Glu Leu Glu Ala Leu Lys Thr Glu Leu Glu Asp Thr Leu 1140 1145 1150 Asp Ser Thr Ala Ala Gln Gln Glu Leu Arg Ser Lys Arg Glu Gln Glu 1155 1160 1165 Val Ser Ile Leu Lys Lys Thr Leu Glu Asp Glu Ala Lys Thr His Glu 1175 1170 1180 Page 245

Ala Gln Ile Gln Glu Met Arg Gln Lys His Ser Gln Ala Val Glu Glu 1195 1190 Leu Ala Glu Gln Leu Glu Gln Thr Lys Arg Val Lys Ala Thr Leu Glu 1205 1210 1215 Lys Ala Lys Gln Thr Leu Glu Asn Glu Arg Gly Glu Leu Ala Asn Glu Val Lys Ala Leu Leu Gln Gly Lys Gly Asp Ser Glu His Lys Arg Lys 1235 1240 1245 Lys Val Glu Ala Gln Leu Gln Glu Leu Gln Val Lys Phe Ser Glu Gly 1250 1255 1260 1255 Glu Arg Val Arg Thr Glu Leu Ala Asp Lys Val Ser Lys Leu Gln Val 1270 Glu Leu Asp Ser Val Thr Gly Leu Leu Asn Gln Ser Asp Ser Lys Ser 1285 Ser Lys Leu Thr Lys Asp Phe Ser Ala Leu Glu Ser Gln Leu Gln Asp 1310 1300 1305 Thr Gln Glu Leu Leu Gln Glu Glu Asn Arg Gln Lys Leu Ser Leu Ser 1320 Thr Lys Leu Lys Gln Met Glu Asp Glu Lys Asn Ser Phe Arg Glu Gln 1335 Leu Glu Glu Glu Glu Glu Ala Lys Arg Asn Leu Glu Lys Gln Ile 1360 1345 1350 1355 Ala Thr Leu His Ala Gln Val Thr Asp Met Lys Lys Met Glu Asp Gly Val Gly Cys Leu Glu Thr Ala Glu Glu Ala Lys Arg Arg Leu Gln 1380 1385 1390 1385 Lys Asp Leu Glu Gly Leu Ser Gln Arg Leu Glu Glu Lys Val Ala Ala 1395 1400 1405 1400 Tyr Asp Lys Leu Glu Lys Thr Lys Thr Arg Leu Gln Gln Glu Leu Asp 1415 Asp Leu Leu Val Asp Leu Asp His Gln Arg Gln Ser Val Ser Asn Leu 1430 1435 Glu Lys Lys Gln Lys Lys Phe Asp Gln Leu Leu Ala Glu Glu Lys Thr 1445 1450 1455 Ile Ser Ala Lys Tyr Ala Glu Glu Arg Asp Arg Ala Glu Ala 1460 1465 1470 Arg Glu Lys Glu Thr Lys Ala Leu Ser Leu Ala Arg Ala Leu Glu Glu 1475 1480 1485 Ala Met Glu Gln Lys Ala Glu Leu Glu Arg Leu Asn Lys Gln Phe Arg 1495 Thr Glu Met Glu Asp Leu Met Ser Ser Lys Asp Asp Val Gly Lys Ser 1505 1510 1515 Page 246

Val His Glu Leu Glu Lys Ser Asn Arg Ala Leu Glu Gln Gln Val Glu 1530 Glu Met Lys Thr Gln Leu Glu Glu Leu Glu Asp Glu Leu Gln Ala Thr 1540 1545 1550 Glu Asp Ala Lys Leu Arg Leu Glu Val Asn Leu Gln Ala Met Lys Ala Gln Phe Glu Arg Asp Leu Gln Gly Arg Asp Glu Gln Ser Glu Glu Lys 1570 1575 1580 Lys Lys Gln Leu Val Arg Gln Val Arg Glu Met Glu Ala Glu Leu Glu 1585 1590 1595 1600 Asp Glu Arg Lys Gln Arg Ser Ile Ala Met Ala Ala Arg Lys Lys Leu Glu Met Asp Leu Lys Asp Leu Glu Ala His Ile Asp Thr Ala Asn Lys Asn Arg Glu Glu Ala Ile Lys Gln Leu Arg Lys Leu Gln Ala Gln Met 1645 1635 1640 Lys Asp Cys Met Arg Asp Val Asp Asp Thr Arg Ala Ser Arg Glu Glu Ile Leu Ala Gln Ala Lys Glu Asn Glu Lys Lys Leu Lys Ser Met Glu 1665 1670 1675 1680 Ala Glu Met Ile Gln Leu Gln Glu Glu Leu Ala Ala Glu Arg Ala 1690 Lys Arg Gln Ala Gln Gln Glu Arg Asp Glu Leu Ala Asp Glu Ile Ala 1700 1705 1710 Asn Ser Ser Gly Lys Gly Ala Leu Ala Leu Glu Glu Lys Arg Arg Leu 1715 1720 1725 Glu Ala Leu Ile Ala Leu Leu Glu Glu Glu Leu Glu Glu Glu Gln Gly 1735 Asn Thr Glu Leu Ile Asn Asp Arg Leu Lys Lys Ala Asn Leu Gln Ile 1745 1750 1755 1760 1750 Asp Gln Ile Asn Thr Asp Leu Asn Leu Glu Arg Ser His Ala Gln Lys 1765 1770 1775 Asn Glu Asn Ala Arg Gln Gln Leu Glu Arg Gln Asn Lys Glu Leu Lys 1785 Ala Lys Leu Gln Glu Met Glu Ser Ala Val Lys Ser Lys Tyr Lys Ala 1795 1800 1805 1800 Ser Ile Ala Ala Leu Glu Ala Lys Ile Ala Gln Leu Glu Glu Gln Leu 1815 Asp Asn Glu Thr Lys Glu Arg Gln Ala Ala Ser Lys Gln Val Arg Arg 1825 1830 1835 Ala Glu Lys Lys Leu Lys Asp Val Leu Leu Gln Val Glu Asp Glu Arg 1845 1850 1855 Page 247

Arg Asn Ala Glu Gln Phe Lys Asp Gln Ala Asp Lys Ala Ser Thr Arg 1860

Leu Lys Gln Leu Lys Arg Gln Leu Glu Glu Ala Glu Glu Ala Gln 1885

Arg Ala Asn Ala Ser Arg Arg Lys Leu Gln Arg Glu Leu Glu Asp Ala 1890

Thr Glu Thr Ala Asp Ala Met Asn Arg Glu Val Ser Ser Leu Lys Asn 1905

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<400> 106

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Trp Val Pro Ser Glu Lys Ser Gly Phe Glu Ala Ala Ser Leu Lys Glu 35 40 45

Glu Val Gly Asp Glu Ala Ile Val Glu Leu Ala Glu Asn Gly Lys Lys
50 55 60

Val Lys Val Asn Lys Asp Asp Ile Gln Lys Met Asn Pro Pro Lys Phe 65 70 75 80

Ser Lys Val Glu Asp Met Ala Glu Leu Thr Cys Leu Asn Glu Ala Ser 85 90 95

Val Leu His Asn Leu Lys Glu Arg Tyr Tyr Ser Gly Leu Ile Tyr Thr 100 105 110

Tyr Ser Gly Leu Phe Cys Val Val Ile Asn Pro Tyr Lys Asn Leu Pro 115 120 125

Ile Tyr Ser Glu Glu Ile Val Glu Met Tyr Lys Gly Lys Lys Arg His 130 140

Glu Met Pro Pro His Ile Tyr Ala Ile Thr Asp Thr Ala Tyr Arg Ser 145 150 155 160

Met Met Gln Asp Arg Glu Asp Gln Ser Ile Leu Cys Thr Gly Glu Ser 165 170 175

Gly Ala Gly Lys Thr Glu Asn Thr Lys Lys Val Ile Gln Tyr Leu Ala Page 248 His Val Ala Ser Ser His Lys Ser Lys Lys Asp Gln Gly Glu Leu Glu 195 200 205 Arg Gln Leu Leu Gln Ala Asn Pro Ile Leu Glu Ala Phe Gly Asn Ala 210 215 220 Lys Thr Val Lys Asn Asp Asn Ser Ser Arg Phe Gly Lys Phe Ile Arg 225 230 235 240 Ile Asn Phe Asp Val Asn Gly Tyr Ile Val Gly Ala Asn Ile Glu Thr 245 250 255 Tyr Leu Leu Glu Lys Ser Arg Ala Ile Arg Gln Ala Lys Glu Glu Arg 260 265 270 Thr Phe His Ile Phe Tyr Tyr Leu Leu Ser Gly Ala Gly Glu His Leu 275 280 285 Thr Asp Leu Leu Glu Pro Tyr Asn Lys Tyr Arg Phe Leu Ser 290 295 300 Asn Gly His Val Thr Ile Pro Gly Gln Gln Asp Lys Asp Met Phe Gln 305 310 315 320 Glu Thr Met Glu Ala Met Arg Ile Met Gly Ile Pro Asp Glu Glu Gln 325 330 335 Ile Gly Leu Leu Lys Val Ile Ser Gly Val Leu Gln Leu Gly Asn Ile 340 345 350 Val Phe Lys Lys Glu Arg Asn Thr Asp Gln Ala Ser Met Pro Asp Asn 355 360 365 Thr Ala Ala Gln Lys Val Ser His Leu Leu Gly Ile Asn Val Thr Asp 370 380 Phe Thr Arg Gly Ile Leu Thr Pro Arg Ile Lys Val Gly Arg Asp Tyr 385 390 395 400 Val Gln Lys Ala Gln Thr Lys Glu Gln Ala Asp Phe Ala Ile Glu Ala 405 410 415 Leu Ala Lys Ala Thr Tyr Glu Gln Met Phe Arg Trp Leu Val Met Arg 420 425 430 Ile Asn Lys Ala Leu Asp Lys Thr Lys Arg Gln Gly Ala Ser Phe Ile 435 440 445 Gly Ile Leu Asp Ile Ala Gly Phe Glu Ile Phe Glu Leu Asn Ser Phe 450 460 Glu Gln Leu Cys Ile Asn Tyr Thr Asn Glu Lys Leu Gln Gln Leu Phe 465 470 475 480 Asn His Thr Met Phe Ile Leu Glu Glu Glu Glu Tyr Gln Asn Glu Gly
485 490 495 Ile Glu Trp Asn Phe Ile Asp Phe Gly Leu Asp Leu Gln Pro Cys Ile 500 505 510Asp Leu Ile Glu Lys Pro Ala Gly Pro Pro Gly Ile Leu Ala Leu Leu Page 249

Asp Glu Glu Cys Trp Phe Pro Lys Ala Thr Asp Lys Ser Phe Val Glu 530 540 Lys Val Val Gln Glu Gln Gly Thr His Pro Lys Phe Gln Lys Pro Lys 545 550 555 560 Gln Leu Lys Asp Lys Ala Asp Phe Cys Ile Ile His Tyr Ala Gly Lys 565 570 575 Val Asp Tyr Lys Ala Asp Glu Trp Leu Met Lys Asn Met Asp Pro Leu 580 585 590 Asn Asp Asn Ile Ala Thr Leu Leu His Gln Ser Ser Asp Lys Phe Val 595 600 605 Ser Glu Leu Trp Lys Asp Val Asp Arg Ile Val Gly Leu Asp Gln Val 610 620 Ala Gly Met Ser Glu Thr Ala Leu Pro Gly Ala Phe Lys Thr Arg Lys 625 630 635 640 Gly Met Phe Arg Thr Val Gly Gln Leu Tyr Lys Glu Gln Leu Ala Lys 645 650 655 Leu Met Ala Thr Leu Arg Asn Thr Asn Pro Asn Phe Val Arg Cys Ile 660 670 Ile Pro Asn His Glu Lys Lys Ala Gly Lys Leu Asp Pro His Leu Val 675 680 685 Leu Asp Gln Leu Arg Cys Asn Gly Val Leu Glu Gly Ile Arg Ile Cys 690 695 700 Arg Gln Gly Phe Pro Asn Arg Val Val Phe Gln Glu Phe Arg Gln Arg 705 710 715 720 Tyr Glu Ile Leu Thr Pro Asn Ala Ile Pro Lys Gly Phe Met Asp Gly 725 730 735 Lys Gln Ala Cys Val Leu Met Ile Lys Ala Leu Glu Leu Asp Ser Asn 740 745 750 Leu Tyr Arg Ile Gly Gln Ser Lys Val Phe Phe Arg Ala Gly Val Leu 755 760 765 Ala His Leu Glu Glu Glu Arg Asp Leu Lys Ile Thr Asp Val Ile Ile 770 775 780 Gly Phe Gln Ala Cys Cys Arg Gly Tyr Leu Ala Arg Lys Ala Phe Ala 785 790 795 800 Lys Arg Gln Gln Gln Leu Thr Ala Met Lys Val Leu Gln Arg Asn Cys 805 810 815 Ala Ala Tyr Leu Lys Leu Arg Asn Trp Gln Trp Trp Arg Leu Phe Thr 820 830 Lys Val Lys Pro Leu Leu Gln Val Ser Arg Gln Glu Glu Met Met 835 840 845 Ala Lys Glu Glu Glu Leu Ile Lys Val Lys Glu Lys Gln Leu Ala Ala

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Glu Asn Arg Leu Ser Glu Met Glu Thr Phe Gln Ala Gln Leu Met Ala 870 Glu Lys Met Gln Leu Gln Glu Gln Leu Gln Ala Glu Ala Glu Leu Cys 885 890 895 Ala Glu Ala Glu Glu Ile Arg Ala Arg Leu Thr Ala Lys Lys Gln Glu 900 905 910 Leu Glu Glu Ile Cys His Asp Leu Glu Ala Arg Val Glu Glu Glu 915 920 925 Glu Arg Cys Gln His Leu Gln Ala Glu Lys Lys Met Gln Gln Asn 930 935 940 Ile Gln Glu Leu Glu Glu Gln Leu Glu Glu Glu Ser Ala Arg Gln 945 950 955 960 Lys Leu Gln Leu Glu Lys Val Thr Thr Glu Ala Lys Leu Lys Lys Leu 965 970 975 Glu Glu Asp Val Ile Val Leu Glu Asp Gln Asn Leu Lys Leu Ala Lys 980 985 990 Glu Lys Lys Leu Leu Glu Asp Arg Met Ser Glu Phe Thr Thr Asn Leu  $995 \hspace{1.5cm} 1000 \hspace{1.5cm} 1005$ Thr Glu Glu Glu Lys Ser Lys Ser Leu Ala Lys Leu Lys Asn Lys 1010 1015 1020 His Glu Ala Met Ile Thr Asp Leu Glu Glu Arg Leu Arg Arg Glu Glu 1025 1030 1035 1040 1025 Lys Gln Arg Gln Glu Leu Glu Lys Thr Arg Arg Lys Leu Glu Gly Asp 1045 1050 1055 Ser Ser Asp Leu His Asp Gln Ile Ala Glu Leu Gln Ala Gln Ile Ala 1065 Glu Leu Lys Ile Gln Leu Ser Lys Lys Glu Glu Leu Gln Ala Ala 1075 1080 1085 Leu Ala Arg Val Glu Glu Glu Ala Ala Gln Lys Asn Met Ala Leu Lys 1090 1095 1100 1090 Lys Ile Arg Glu Leu Glu Ser Gln Ile Thr Glu Leu Gln Glu Asp Leu 1115 1110 Glu Ser Glu Arg Ala Ser Arg Asn Lys Ala Glu Lys Gln Lys Arg Asp 1125 1130 1135 Leu Gly Glu Glu Leu Glu Ala Leu Lys Thr Glu Leu Glu Asp Thr Leu 1140 1145 1150 1140 Asp Ser Thr Ala Ala Gln Gln Glu Leu Arg Ser Lys Arg Glu Gln Glu 1160 Val Thr Val Leu Lys Lys Thr Leu Glu Asp Glu Ala Lys Thr His Glu 1170 1175 1180 Ala Gln Ile Gln Glu Met Arg Gln Lys His Ser Gln Ala Ile Glu Glu

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Leu Ala Glu Gln Leu Glu Gln Thr Lys Arg Val Lys Ala Asn Leu Glu 1205 1210 1215

Lys Ala Lys Gln Ala Leu Glu Ser Glu Arg Ala Glu Leu Ser Asn Glu 1220 1225 1230

Val Lys Val Leu Leu Gln Gly Lys Gly Asp Ala Glu His Lys Arg Lys 1235 1240 1245

Lys Val Asp Ala Gln Leu Gln Glu Leu Gln Val Lys Phe Thr Glu Gly 1255

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Glu Leu Asp Asn Val Thr Gly Leu Leu Asn Gln Ser Asp Ser Lys Ser 1285 1290 1295

Ile Lys Leu Ala Lys Asp Phe Ser Ala Leu Glu Ser Gln Leu Gln Asp 1305

Thr Gln Glu Leu Leu Gln Glu Glu Thr Arg Leu Lys Leu Ser Phe Ser 1315 1320 1325

Thr Lys Leu Lys Gln Thr Glu Asp Glu Lys Asn Ala Leu Lys Glu Gln 1335

Leu Glu Glu Glu Glu Ala Lys Arg Asn Leu Glu Lys Gln Ile Ser 1350 1355

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Leu Gly Cys Leu Glu Ile Ala Glu Glu Ala Lys Lys Lys Leu Gln Lys 1385

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Asp Lys Leu Glu Lys Thr Lys Thr Arg Leu Gln Gln Glu Leu Asp Asp 1410 1415 1420

Ile Ala Val Asp Leu Asp His Gln Arg Gln Thr Val Ser Asn Leu Glu 1430 1435 1425

Lys Lys Gln Lys Lys Phe Asp Gln Leu Leu Ala Glu Glu Lys Asn Ile 1445 1450 1455

Ser Ala Lys Tyr Ala Glu Glu Arg Asp Arg Ala Glu Ala Glu Ala Arg 1460 1465 1470

Glu Lys Glu Thr Lys Ala Leu Ser Leu Ala Arg Ala Leu Glu Glu Ala 1480

Ile Glu Gln Lys Ala Glu Leu Glu Arg Val Asn Lys Gln Phe Arg Thr 1495

Glu Met Glu Asp Leu Met Ser Ser Lys Asp Asp Val Gly Lys Ser Val 1505 1510 1520

His Glu Leu Glu Lys Ala Lys Arg Ala Leu Glu Gln Gln Val Glu Glu Page 252

Met Lys Thr Gln Leu Glu Glu Leu Glu Asp Glu Leu Gln Ala Thr Glu 1545 Asp Ala Lys Leu Arg Leu Glu Val Asn Gln Gln Ala Met Lys Ala Gln 1555 1560 1565 Phe Asp Arg Asp Leu Leu Gly Arg Asp Glu Gln Asn Glu Glu Lys Arg 1570 1575 1580 Lys Gln Leu Ile Arg Gln Val Arg Glu Met Glu Val Glu Leu Glu Asp 1585 1590 1595 1600 Glu Arg Lys Gln Arg Ser Ile Ala Val Ala Ala Arg Lys Lys Leu Glu 1605 1610 1615 Leu Asp Leu Lys Asp Leu Glu Ser His Ile Asp Thr Ala Asn Lys Asn 1625 Arg Asp Glu Ala Ile Lys His Val Arg Lys Leu Gln Ala Gln Met Lys Asp Tyr Met Arg Glu Leu Glu Asp Thr Arg Thr Ser Arg Glu Glu Ile 1650 1660 Leu Ala Gln Ala Lys Glu Asn Glu Lys Lys Leu Lys Ser Met Glu Ala 1665 1670 1675 1680 Glu Met Ile Gln Leu Gln Glu Glu Leu Ala Ala Glu Arg Ala Lys Arg Gln Ala Gln Glu Arg Asp Glu Leu Ala Asp Glu Ile Ala Asn 1700 1705 1710 Ser Ser Gly Lys Gly Ala Leu Ala Met Glu Glu Lys Arg Arg Leu Glu 1715 1720 1725 Ala Arg Ile Ala Gln Leu Glu Glu Glu Leu Glu Glu Glu Gln Gly Asn Thr Glu Ile Ile Asn Asp Arg Leu Lys Lys Ala Asn Leu Gln Ile Asp 1745 1750 1755 1760 Gln Met Asn Ala Asp Leu Asn Ala Glu Arg Ser Asn Ala Gln Lys Asn 1765 1770 1775 Glu Asn Ala Arg Gln Gln Met Glu Arg Gln Asn Lys Glu Leu Lys Leu 1780 1785 1790 Lys Leu Gln Glu Met Glu Ser Ala Val Lys Ser Lys Tyr Lys Ala Thr 1795 1800 1805 Ile Thr Ala Leu Glu Ala Lys Ile Val Gln Leu Glu Gln Leu Asp 1810 1815 1820 Met Glu Thr Lys Glu Arg Gln Ala Ala Ser Lys Gln Val Arg Arg Ala 1825 1830 1835 1840 Glu Lys Lys Leu Lys Asp Ile Leu Leu Gln Val Asp Asp Glu Arg Arg 1845 1850 1855 Asn Ala Glu Gln Phe Lys Asp Gln Ala Asp Lys Ala Asn Met Arg Leu

1870

Lys Gln Leu Lys Arg Gln Leu Glu Glu Ala Glu Glu Glu Ala Gln Arg 1875 1880 1885

Ala Asn Val Arg Arg Lys Leu Gln Arg Glu Leu Asp Asp Ala Thr Glu 1890 1895 1900

Thr Ala Asp Ala Met Asn Arg Glu Val Ser Ser Leu Lys Ser Lys Leu 1905 1910 1915 1920

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50 55 60

Val Lys Val Asn Lys Asp Asp Ile Gln Lys Met Asn Pro Pro Lys Phe 65 70 75 80

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Glu Met Pro Pro His Ile Tyr Ala Ile Thr Asp Thr Ala Tyr Arg Ser 145 150 155 160

Met Met Gln Asp Arg Glu Asp Gln Ser Ile Leu Cys Thr Gly Glu Ser 165 170 175

Gly Ala Gly Lys Thr Glu Asn Thr Lys Lys Val Ile Gln Tyr Leu Ala 180 185 190 CURA2221.APP
His Val Ala Ser Ser His Lys Ser Lys Lys Asp Gln Gly Glu Leu Glu
195 200 205 Arg Gln Leu Leu Gln Ala Asn Pro Ile Leu Glu Ala Phe Gly Asn Ala 210 215 220 Lys Thr Val Lys Asn Asp Asn Ser Ser Arg Phe Gly Lys Phe Ile Arg 225 230 235 240 Ile Asn Phe Asp Val Asn Gly Tyr Ile Val Gly Ala Asn Ile Glu Thr 245 250 255 Tyr Leu Leu Glu Lys Ser Arg Ala Ile Arg Gln Ala Lys Glu Glu Arg 260 265 270 Thr Phe His Ile Phe Tyr Tyr Leu Leu Ser Gly Ala Gly Glu His Leu 275 280 285 Lys Thr Asp Leu Leu Glu Pro Tyr Gly Lys Tyr Arg Phe Leu Ser 290 295 300 Asn Gly His Val Thr Ile Pro Gly Gln Gln Asp Lys Asp Met Phe Gln 305 310 315 320 Glu Thr Met Glu Ala Met Arg Ile Met Gly Ile Pro Asp Glu Gln 325 330 335 Ile Gly Leu Leu Lys Val Ile Ser Gly Val Leu Gln Leu Gly Asn Ile 340 345 350 Val Phe Lys Lys Glu Arg Asn Thr Asp Gln Ala Ser Met Pro Asp Asn 355 360 365 Thr Ala Ala Gln Lys Val Ser His Leu Leu Gly Ile Asn Val Thr Asp 370 375 380 Phe Thr Arg Gly Ile Leu Thr Pro Arg Ile Lys Val Gly Arg Asp Tyr 385 390 395 400 Val Gln Lys Ala Gln Thr Lys Glu Gln Ala Asp Phe Ala Ile Glu Ala 405 410 415 Leu Ala Lys Ala Thr Tyr Glu Gln Met Phe Arg Trp Leu Val Met Arg 420 425 430 Ile Asn Lys Ala Leu Asp Lys Thr Lys Arg Gln Gly Ala Ser Phe Ile 435 440 445 Gly Ile Leu Asp Ile Ala Gly Phe Glu Ile Phe Glu Leu Asn Ser Phe 450 455 460 Glu Gln Leu Cys Ile Asn Tyr Thr Asn Glu Lys Leu Gln Gln Leu Phe 465 470 475 480 Asn His Thr Met Phe Ile Glu Gln Glu Glu Tyr Gln Arg Glu Gly Ile 485 490 495 Glu Trp Asn Phe Ile Asp Phe Gly Leu Asp Leu Gln Pro Cys Ile Asp 500 505 510 Leu Ile Glu Arg Pro Ala Asn Pro Pro Gly Val Leu Ala Leu Leu Asp 515 520 525 Glu Glu Cys Trp Phe Pro Lys Ala Thr Asp Lys Thr Phe Val Glu Lys 530 535 540 Leu Val Gln Glu Gln Gly Thr His Ser Lys Phe Gln Lys Pro Arg Gln 545 550 555 560 Leu Lys Asp Lys Ala Asp Phe Cys Ile Ile His Tyr Ala Gly Lys Val 565 570 575 Asp Tyr Lys Gly Asp Glu Trp Leu Met Lys Asn Met Asp Pro Leu Asn 580 585 590 Asp Asn Val Ala Thr Leu Leu His Gln Ser Ser Asp Lys Phe Val Ala 595 600 605 Glu Leu Trp Lys Asp Val Asp Arg Ile Val Gly Leu Asp Gln Val Thr 610 615 620 Gly Ile Thr Glu Thr Ala Phe Gly Ser Ala Tyr Lys Thr Lys Lys Gly 625 630 635 640 Met Phe Arg Thr Val Gly Gln Leu Tyr Lys Glu Ser Leu Thr Lys Leu 645 650 655 Met Ala Thr Leu Arg Asn Thr Asn Pro Asn Phe Val Arg Cys Ile Ile 660 665 670 Pro Asn His Glu Lys Arg Ala Gly Lys Leu Asp Pro His Leu Val Leu 675 680 685 Asp Gln Leu Arg Cys Asn Gly Val Leu Glu Gly Ile Arg Ile Cys Arg 690 695 700 Gln Gly Phe Pro Asn Arg Ile Val Phe Gln Glu Phe Arg Gln Arg Tyr 705 710 715 720 Glu Ile Leu Thr Pro Asn Ala Ile Pro Lys Gly Phe Met Asp Gly Lys 725 730 735 Gln Ala Cys Glu Arg Met Ile Arg Ala Leu Glu Leu Asp Pro Asn Leu 740 745 750 Tyr Arg Ile Gly Gln Ser Lys Ile Phe Phe Arg Ala Gly Val Leu Ala 755 760 765 His Leu Glu Glu Glu Arg Asp Leu Lys Ile Thr Asp Ile Ile Ile Phe 770 775 780 Phe Gln Ala Val Cys Arg Gly Tyr Leu Ala Arg Lys Ala Phe Ala Lys 785 790 795 800 Lys Gln Gln Leu Ser Ala Leu Lys Ile Leu Gln Arg Asn Cys Ala 805 810 815 Ala Tyr Leu Lys Leu Arg His Trp Gln Trp Trp Arg Val Phe Thr Lys 820 825 830 Val Lys Pro Leu Leu Gln Val Thr Arg Gln Glu Glu Leu Gln Ala 835 840 845 Lys Asp Glu Glu Leu Met Lys Lys Val Glu Lys Gln Thr Lys Val Glu 850 855 860

CURA2221.APP Ala Glu Leu Glu Glu Met Glu Arg Lys His Gln Gln Leu Leu Glu Glu Lys Asn Ile Leu Ala Glu Gln Leu Gln Ala Glu Thr Glu Leu Phe Ala 885 890 895 Glu Ala Glu Glu Met Arg Ala Arg Leu Ala Ala Lys Lys Gln Glu Leu 900 905 910 Glu Glu Ile Leu His Asp Leu Glu Ser Arg Val Glu Glu Glu Glu 915 920 925 Arg Asn Gln Ile Leu Gln Asn Glu Lys Lys Glu Gln Gly His Lys 930 935 940 Asn Asp Leu Glu Glu Gln Leu Asp Glu Met Glu Ser Ala Arg Gln Lys 945 950 955 960 Leu Gln Leu Glu Lys Val Thr Thr Glu Ala Lys Leu Lys Lys Leu Glu 965 970 975 Glu Glu Gln Ile Ile Leu Glu Asp Gln Asn Cys Lys Leu Ala Lys Glu 980 985 990 Lys Lys Leu Leu Glu Asp Arg Ile Ala Glu Phe Thr Thr Asn Leu Thr 995 1000 1005 Glu Glu Glu Lys Ser Lys Ser Leu Ala Lys Leu Lys Asn Lys His 1010 1015 1020 Glu Ala Met Ile Thr Asp Leu Glu Glu Arg Leu Arg Arg Glu Glu Lys 1025 1030 1035 1040 Gln Arg Gln Glu Leu Glu Lys Thr Arg Arg Lys Leu Glu Gly Asp Ser 1045 1050 1055 Thr Asp Leu Ser Asp Gln Ile Ala Glu Leu Gln Ala Gln Ile Ala Glu 1065 Leu Lys Met Gln Leu Ala Lys Lys Glu Glu Glu Leu Gln Ala Ala Leu 1075 1080 1085 Ala Arg Val Glu Glu Ala Ala Gln Lys Asn Met Ala Leu Lys Lys Ile Arg Glu Leu Glu Ser Gln Ile Ser Glu Leu Gln Glu Asp Leu Glu 1115 Ser Glu Arg Ala Ser Arg Asn Lys Ala Glu Lys Gln Lys Arg Asp Leu 1125 1130 1135 Gly Glu Glu Leu Glu Ala Leu Lys Thr Glu Leu Glu Asp Leu Thr Asp Ser Thr Ala Ala Gln Gln Glu Leu Arg Ser Lys Arg Glu Gln Glu Val Asn Ile Leu Lys Lys Thr Leu Glu Glu Glu Ala Lys Thr His Glu Ala 1170 1175 1180

Gln Ile Gln Glu Met Arg Gln Lys His Ser Gln Ala Val Glu Glu Leu 1185 1190 1195 1200

1185

Ala Glu Gln Leu Glu Gln Thr Lys Arg Lys Val Ala Asn Leu Glu Lys 1205 1210 1215

Ala Lys Gln Thr Leu Glu Asn Glu Arg Gly Glu Leu Ala Asn Glu Val 1220 1225 1230

Lys Val Leu Leu Gln Gly Gly Arg Asp Ser Glu His Lys Arg Lys Lys 1235 1240 1245

Val Glu Ala Gln Leu Gln Glu Leu Gln Val Lys Phe Asn Glu Gly Glu 1250 1255 1260

Leu Asp Asn Val Thr Gly Leu Leu Ser Gln Ser Asp Ser Lys Ser Ser 1285 1290 1295

Lys Leu Thr Lys Asp Phe Ser Ala Leu Glu Ser Gln Leu Gln Asp Thr 1300 1305 1310

Gln Glu Leu Leu Gln Glu Glu Asn Arg Gln Lys Leu Ser Leu Ser Thr 1315 1320 1325

Lys Leu Lys Gln Val Glu Asp Glu Lys Asn Ser Phe Arg Glu Gln Leu 1330 1335 1340

Glu Glu Glu Glu Glu Ala Lys His Asn Leu Glu Lys Gln Ile Ala 1345 1350 1355 1360

Thr Leu His Ala Gln Val Ala Asp Met Lys Lys Met Glu Asp Ser 1365 1370 1375

Val Gly Cys Leu Glu Thr Ala Glu Glu Val Lys Arg Lys Leu Gln Lys 1380 1385 1390

Asp Leu Glu Gly Leu Ser Gln Arg His Glu Glu Lys Val Ala Ala Tyr 1395 1400 1405

Asp Lys Leu Glu Lys Thr Lys Thr Arg Leu Gln Gln Glu Leu Asp Asp 1410 1415 1420

Leu Leu Val Asp Leu Asp His Gln Arg Gln Ser Ala Cys Asn Leu Glu 1425 1430 1435 1440

Lys Lys Gln Lys Lys Phe Asp Gln Leu Leu Ala Glu Glu Ile Thr Lys  $1445 \hspace{1cm} 1450 \hspace{1cm} 1455$ 

Ser Ala Lys Tyr Ala Glu Glu Arg Ala Arg Asp Ala Glu Glu Arg Ala 1460 1465 1470

Glu Lys Ala Thr Lys Glu Leu Ser Leu Ala Arg Ala Glu Leu Glu Ala 1475 1480 1485

Met Glu Gln Lys Ala Glu Phe Leu Arg Lys Asn Leu Gln Glu Met Thr 1490 1495 1500

Glu Arg Leu Asp Glu Met Ser Ser Lys Val Asp Asp Ala Lys Ser Val 1505 1510 1515 1520

Leu Glu His Glu Lys Ser Lys Leu Gly Arg Glu Gln Gln Val Met Glu 1525 1530 1535

Cl	JRA	.22	221	. A	PP

Glu Lys Thr Gln Leu Leu Glu Glu Glu Asp Glu Leu Ala Gln Thr Glu 1540 1545 1550

Asp Ala Lys Leu Arg Leu Glu Val Asn Leu Gln Ala Met Lys Ala Gln 1555 1560 1565

Phe Glu Arg Asp Leu Gln Gly Arg Gln Asp Asp Ser Glu Glu Lys Gln 1570 1575 1580

Lys Lys Leu Val Arg Gln Val Arg Glu Met Glu Ala Glu Leu Glu Asp 1585 1590 1595 1600

Gln Arg Lys Glu Met Ser Arg Ala Arg Ala Ala Val Lys Lys Leu Glu 1605 1610 1615

Met Asp Leu Lys Asp Leu Glu Ala His Ile Asp Ser Ala Asn Lys Asn 1620 1625 1630

Arg Asp Glu Ala Lys Ile Gln Leu Arg Asn Leu Gln Ala Gln Met Lys 1635 1640 1645

Asp Cys Met Arg Glu Leu Asp Asp Thr Arg Ala Ser Arg Glu Glu Ile 1650 1655 1660

Ala Leu Gln Ala Lys Glu Asn Glu Lys Lys Leu Lys Ser Met Glu Ala 1665 1670 1675 1680

Glu Met Ile Gln Leu Gln Glu Glu Leu Ala Ala Glu Arg Ala Lys 1685 1690 1695

Arg Gln Ala Gln Glu Arg Asp Glu Leu Ala Asp Glu Ile Ser Asn 1700 1705 1710

Ala Ser Gly Lys Ala Gly Leu Ala Lys Glu Glu Leu Arg Arg Leu Glu 1715 1720 1725

Ala Arg Ile Ala Gln Leu Glu Glu Glu Glu Glu Gln Gly Asn 1730 1740

Thr Glu Leu Ile Asn Asp Arg Leu Lys Lys Ala Asn Leu Gln Ile Asp 1745 1750 1760

Gln Ile Asn Ala Asp Leu Asn Leu Glu Arg Gly His Ala Gln Lys Asn 1765 1770 1775

Glu Asn Ala Arg Gln Gln Leu Glu Arg Gln Asn Lys Glu Leu Lys Val 1780 1785 1790

Lys Leu Gln Glu Met Glu Gly Thr Val Lys Ser Lys Tyr Lys Ala Ser 1795 1800 1805

Ile Thr Ala Leu Glu Ala Lys Ile Ala Gln Leu Glu Gln Leu Asp 1810 1815 1820

Asn Glu Thr Lys Glu Arg Gln Ala Ala Cys Lys Gln Val Arg Arg Thr 1825 1830 1835 1840

Glu Lys Lys Leu Lys Asp Val Leu Leu Gln Val Asp Asp Glu Arg Arg 1845 1850 1855

Asn Ala Glu Gln Tyr Lys Asp Gln Ala Asp Lys Ala Ser Thr Arg Leu 1860 1865 1870

Lys Gln Leu Lys Arg Gln Leu Glu Glu Ala Glu Glu Glu Ala Gln Arg 1875 1880 1885

Ala Asn Ala Ser Arg Arg Lys Leu Gln Arg Glu Leu Glu Asp Ala Thr 1890 1895 1900

Glu Thr Ala Asp Ala Met Asn Arg Glu Val Ser Ser Leu Lys Asn Lys 1905 1910 1915 1920

Leu Arg Arg Gly Asp Leu Pro Phe Val Val Thr Arg Arg Leu Val Arg 1925 1930 1935

Lys Gly Thr Leu Glu Leu Ser Asp Asp Asp Glu Ser Lys Ala Ser 1940 1945 1950

Leu Ile Asn Glu Thr Gln Pro Pro Gln Cys Leu Asp Gln Gln Leu Asp 1955 1960 1965

Gln Leu Phe His Trp Pro Val Asn Ala Gly Cys Val Cys Gly Trp Gly 1970 1975 1980

Val Glu Gln Thr Gln Gly Glu Glu Ala Val His Lys Cys Arg Thr 1985 1990 1995

<210> 108

<211> 734

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Myosin Head (motor domain) sequence

<400> 108

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His Asn Leu Lys Lys Arg Tyr Lys Ser Asp Leu Ile Tyr Thr Tyr Ser . 20 25 30

Gly Leu Val Leu Val Ser Val Asn Pro Tyr Lys Arg Leu Pro Gln Ile 35 40 45

Tyr Thr Glu Glu Ile Ile Ala Lys Tyr Arg Gly Lys Arg Arg Tyr Glu 50 55 60

Leu Pro Pro His Ile Phe Ala Ile Ala Asp Glu Ala Tyr Arg Ser Met 65 70 75 80

Leu Ser Asp Lys Glu Asn Gln Ser Ile Leu Ile Ser Gly Glu Ser Gly 85 90 95

Ala Gly Lys Thr Glu Asn Thr Lys Lys Val Met Gln Tyr Leu Ala Ala 100 105 110

Val Ser Gly Gly Asn Ser Gly Asn Gly Glu Glu Val Pro Ser Val Lys 115 120 125

Val Gly Arg Val Glu Asp Gln Ile Leu Gln Ser Asn Pro Ile Leu Glu 130 135 140

Ala Phe Gly Asn Ala Lys Thr Thr Arg Asn Asn Asn Ser Ser Arg Phe Page 260

150 160 145 Gly Lys Tyr Ile Glu Ile Gln Phe Asp Lys Thr Gly Lys Ile Val Gly 165 170 175 Ala Lys Ile Glu Asn Tyr Leu Leu Glu Lys Ser Arg Val Val Tyr Gln 180 185 190 Thr Glu Gly Glu Arg Asn Phe His Ile Phe Tyr Gln Leu Leu Ala Gly 195 200 205 Ala Ser Gln Gln Asn Leu Lys Lys Glu Leu Lys Leu Thr Asn Asp Pro 210 215 220 Glu Asp Tyr His Tyr Leu Asn Gln Gly Gly Glu Val Lys Pro Cys Tyr 225 230 235 240 Thr Val Asp Gly Ile Asp Asp Ser Glu Gly Asn Val Glu Glu Phe Lys 245 250 255 Glu Thr Arg Lys Ala Met Asp Ile Leu Gly Phe Thr Asp Glu Glu Gln 260 265 270 Arg Ser Ile Phe Arg Ile Val Ala Ala Ile Leu His Leu Gly Asn Ile 275 280 285 Phe Lys Gln Arg Arg Lys Glu Glu Ala Ala Ile Pro Asp Asp Asn 290 295 300 Asn Ala Asp Thr Lys Ala Leu Glu Lys Ala Ala Glu Leu Leu Gly Val 305 310 315 320 Asp Ala Thr Glu Leu Glu Lys Ala Leu Leu Ser Arg Arg Ile Lys Thr 325 330 335 Gly Thr Glu Gly Arg Lys Ser Thr Val Thr Lys Pro Gln Asn Val Glu 340 345 350 Gln Ala Ser Tyr Ala Arg Asp Ala Leu Ala Lys Ala Leu Tyr Ser Arg 355 360 365 Leu Phe Asp Trp Ile Val Asn Arg Ile Asn Lys Thr Leu Asp Phe Lys 370 375 380 Ala Lys Glu Gly Gln Asp Ala Ser Phe Ile Gly Val Leu Asp Ile Tyr 385 390 400 Gly Phe Glu Ile Phe Glu Lys Asn Ser Phe Glu Gln Leu Cys Ile Asn 405 410 415 Tyr Val Asn Glu Lys Leu Gln Gln Phe Phe Asn His His Met Phe Lys 420 425 430 Leu Glu Gln Glu Glu Tyr Lys Arg Glu Gly Ile Glu Trp Thr Phe Ile 435 440 445 Asp Phe Gly Asp Asn Leu Gln Pro Cys Ile Asp Leu Ile Glu Lys Lys Ser Pro Pro Gly Ile Leu Ser Leu Leu Asp Glu Glu Cys Leu Phe Pro 465 470 475 480

Lys Ala Gln Ser Gly Thr Asp Gln Thr Phe Leu Asp Lys Leu Tyr Ser

Thr Phe Ser Lys His Pro Ala His Phe Glu Lys Phe Ser Pro Arg Phe 500 505 510 Arg Gln Lys Lys Ser Gly Ala His Phe Ile Ile Lys His Tyr Ala Gly 515 520 525 Val Glu Tyr Asn Val Glu Gly Phe Leu Glu Lys Asn Lys Asp Pro 530 540 Leu Phe Asp Asp Leu Ile Ser Leu Leu Lys Ser Ser Ser Asn Pro Leu 545 555 560 Leu Ala Glu Leu Phe Pro Asp Glu Glu Thr Leu Ala Gly Pro Phe Glu 565 570 575 Ala Asp Pro Ser Ser Leu Ser Lys Lys Arg Lys Ser Gly Ser Lys Asn 580 585 590 Lys Ser Thr Gly Lys Lys Thr Lys Lys Ser Asn Phe Ile Thr Val Gly 595 600 605 Ala Gln Phe Lys Glu Ser Leu Asn Glu Leu Met Lys Thr Leu Ser Ser 610 615 620 Thr Asn Leu Pro His Phe Val Arg Cys Ile Lys Pro Asn Glu Lys Lys 625 630 635 640 Lys Ala Gly Val Phe Asp Ala Ser Leu Val Leu His Gln Leu Arg Cys 645 650 655 Leu Gly Val Leu Glu Gly Ile Arg Ile Arg Arg Ala Gly Phe Pro Asn 660 665 670 Arg Ile Thr Phe Asp Glu Phe Leu Gln Arg Tyr Arg Ile Leu Ala Pro 675 680 685 Lys Thr Trp Pro Lys Trp Ser Gly Asp Ala Lys Lys Gly Glu Lys Asn 690 695 700 Glu Ile Val Ala Cys Glu Lys Leu Leu Gln Ser Leu Asn Leu Asp Lys 705 710 715 720 Gly Glu Glu Tyr Arg Phe Gly Lys Thr Lys Ile Phe Phe Arg 725 730

<210> 109

<211> 175

<212> PRT

<213> Homo sapiens

<400> 109

Met Leu Pro Pro Met Ala Leu Pro Ser Val Ser Trp Met Leu Leu Ser 1 5 10

Cys Leu Met Leu Leu Ser Gln Val Gln Gly Glu Glu Pro Gln Arg Glu 20 25 30

Leu Pro Ser Ala Arg Ile Arg Cys Pro Lys Gly Ser Lys Ala Tyr Gly

Ser His Cys Tyr Ala Leu Phe Leu Ser CURA2221.APP Pro Lys Ser Trp Thr Asp Ala Asp Leu Ala Cys Gln Lys Arg Pro Ser Gly Asn Leu Val Ser Val Leu 80 Ser Gly Ala Glu Gly Ser Phe Val Ser Ser Leu Val Lys Ser Ile Gly Asn Ser Tyr Ser Tyr Val Trp Ile Gly Leu His Asp Pro Thr Gln Gly 110 Thr Glu Pro Asn Gly Glu Gly Trp Glu Trp Ser Ser Ser Asp Val Met Asn Tyr Phe Ala Trp Glu Arg Asn Pro Ser Thr Ile Ser Ser Pro Gly His Cys Ala Ser Leu Ser Arg Ser Thr Ala Phe Leu Arg Trp Lys Asp 145 Tyr Asn Cys Asn Val Arg Leu Pro Tyr Val Cys Lys Phe Thr Asp 175

<210> 110 <211> 175 <212> PRT

<213> Bos taurus

 4400> 110
 Met Leu Pro Ser Leu Gly Leu Pro Arg Leu Ser Trp Met Leu Leu Ser 15

 Cys Leu Met Leu Leu Ser Gln Ile Gln Gly Glu Asn Ser Gln Lys Glu 25

 Leu Pro Ser Ala Arg Ile Ser Cys Pro Ser Gly Ser Met Ala Tyr Arg 35

 Ser His Cys Tyr Ala Leu Phe 55
 Lys Thr Pro Lys Thr Trp Met Asp Ala 60

 Asp Ile Ala Cys Gln Lys Arg Pro Ser Gly His Leu Val Ser Val Leu 80

 Ser Gly Ala Glu Glu Ser Phe Val Ala Ser Leu Val Arg Asn Asn Leu 95

 Asn Thr Gln Ser Asp Ile Trp Ile Gly Leu His Asp Pro Thr Glu Gly 110

 Ser Glu Ala Asn Ala Gly Gly Trp Glu Trp Ile Ser Asn Asp Val Leu Asn Tyr Val Ala Trp Glu Thr Asp Pro Ala Ala Ile Ser Ser Pro Gly 145

 Tyr Cys Gly Ser Leu Ser Arg Ser Ser Gly Tyr Leu Lys Trp Arg Asp 160

 His Asn Cys Asn Leu Asn Leu Pro Tyr Val Cys Lys Phe Thr Asp 175

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<210> 111
<211> 175
<212> PRT
<213> Rattus norvegicus
<400> 111
Met Leu His Arg Leu Ala Phe Pro Val Met Ser Trp Met Leu Leu Ser
1 10 15
Cys Leu Met Leu Leu Ser Gln Val Gln Gly Glu Asp Ser Pro Lys Lys 20 25 30
Ile Pro Ser Ala Arg Ile Ser Cys Pro Lys Gly Ser Gln Ala Tyr Gly
Ser Tyr Cys Tyr Ala Leu Phe Gln Ile Pro Gln Thr Trp Phe Asp Ala
50 55 60
Glu Leu Ala Cys Gln Lys Arg Pro Glu Gly His Leu Val Ser Val Leu 65 70 75 80
Asn Val Ala Glu Ala Ser Phe Leu Ala Ser Met Val Lys Asn Thr Gly
85 90 95
Asn Ser Tyr Gln Tyr Thr Trp Ile Gly Leu His Asp Pro Thr Leu Gly 100 	 105 	 110
Gly Glu Pro Asn Gly Gly Gly Trp Glu Trp Ser Asn Asn Asp Ile Met
115 120 125
Asn Tyr Val Asn Trp Glu Arg Asn Pro Ser Thr Ala Leu Asp Arg Gly 130 140
Phe Cys Gly Ser Leu Ser Arg Ser Ser Gly Phe Leu Arg Trp Arg Asp 145 150 155 160
Thr Thr Cys Glu Val Lys Leu Pro Tyr Val Cys Lys Phe Thr Gly
<210> 112
<211> 175
<212> PRT
<213> Mus musculus
<400> 112
Met Leu Pro Pro Thr Ala Cys Ser Val Met Ser Trp Met Leu Leu Ser 1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15
Cys Leu Met Leu Leu Ser Gln Val Gln Gly Glu Asp Ser Leu Lys Asn
20 25 30
Ile Pro Ser Ala Arg Ile Ser Cys Pro Lys Gly Ser Gln Ala Tyr Gly 40 45
Ser Tyr Cys Tyr Ala Leu Phe Gln Ile Pro Gln Thr Trp Phe Asp Ala 50 55 60
```

Glu Leu Ala Cys Gln Lys Arg Pro Gly Gly His Leu Val Ser Val Leu 65 70 75 80 Asn Ser Ala Glu Ala Ser Phe Leu Ser Ser Met Val Lys Arg Thr Gly

Asn Ser Tyr Gln Tyr Thr Trp Ile Gly Leu His Asp Pro Thr Leu Gly

Ala Glu Pro Asn Gly Gly Gly Trp Glu Trp Ser Asn Asn Asp Val Met

115 Asn Tyr Phe Asn Trp Glu Arg Asn Pro Ser Thr Ala Leu Asp Arg Ala

Phe Cys Gly Ser Leu Ser Arg Ala Ser Gly Phe Leu Lys Trp Arg Asp

145 Asn Thr Cys Glu Val Lys Leu Pro Tyr Val Cys Lys Phe Thr Gly

175 Asp

2210> 113

2211> 174

2212> PRT

2213> Rattus norvegicus

4400> 113

Met Leu Pro Arg Val Ala Leu Thr Thr Met Ser Trp Met Leu Leu Ser

Ser Leu Met Leu Leu Ser Gln Val Gln Gly Glu Asp Ala Lys Glu Asp

Val Pro Thr Ser Arg Ile Ser Cys Pro Lys Gly Ser Arg Ala Tyr Gly

Ser Leu Met Leu Leu Ser Gln Val Gln Gly Glu Asp Ala Lys Glu Asp Val Pro Thr Ser Arg Ile Ser Cys Pro Lys Gly Ser Arg Ala Tyr Gly Ser Tyr Cys Tyr Ala Leu Phe Ser Val Ser Lys Ser Trp Phe Asp Ala Asp Leu Ala Cys Gln Lys Arg Pro Ser Gly His Leu Val Ser Val Leu 80 Ser Gly Ser Gly Ser Glu Ala Ser Phe Val Ser Ser Leu Ile Lys Ser Ser Gly Asn Ser Gly Gln Asn Val Trp Ile Gly Leu His Asp Pro Thr Leu Gly 110 Gln Glu Pro Asn Arg Gly Gly Trp Glu Trp Ser Asn Ala Asp Val Met Asn Tyr Phe Asn Trp Glu Thr Asn Pro Ser Ser Val Ser Gly Ser Gly Ser His Cys Gly Thr Leu Thr Arg Ala Ser Gly Phe Leu Arg Trp Arg Glu Asn Cys Ile Ser Glu Leu Pro Tyr Val Cys Lys Phe Lys Ala

<210> 114 <211> 125 <212> PRT <213> Artificial Sequence

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       domain sequence
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1 5 10 15
Gly His Ala His Leu Val Ser Ile Gln Ser Ala Glu Glu Gln Ser Phe
20 25 30
Val Val Ala Phe Leu Thr Ser Leu Thr Lys Lys Ser Asn Thr Tyr Ala 45
Trp Ile Gly Leu Thr Asp Ile Asn Thr Glu Gly Thr Trp Val Trp Glu 50 60
Gly Trp Glu Thr Asp Gly Ser Pro Val Asn Tyr Thr Glu Asn Trp Ala
65 70 75 80
Pro Gly Glu Pro Asn Asn Arg Gly Asn His Gly Gly Asn Glu Asp Cys
85
90
95
Val Glu Ile Tyr Thr Asp Thr Asp Phe Leu Ala Gly Lys Trp Asn Asp 100 \hspace{1cm} 105 \hspace{1cm} 110
Glu Pro Cys Asp Ser Lys Leu Pro Tyr Val Cys Glu Phe
115 120 125
<210> 115
<211> 21
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: PCR Primer
       sequence
<400> 115
                                                                             21
ctggttgtag gttgccatgg t
<210> 116
<211> 24
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: PCR Primer
       sequence
<400> 116
cagcttcgtt ggcacaggcc tctc
                                                                             24
<210> 117
<211> 26
<212> DNA
<213> Artificial Sequence
<220>
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# CURA2221.APP <223> Description of Artificial Sequence: PCR Primer sequence <400> 117 26 ccagtataag ctgacctttg acaaag <210> 118 <211> 21 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: PCR Primer sequence <400> 118 21 ctggttgtag gttgccatgg t <210> 119 <211> 24 <212> DNA <213> Artificial Sequence <223> Description of Artificial Sequence: PCR Primer sequence <400> 119 24 cagcttcgtt ggcacaggcc tctc <210> 120 <211> 26 <212> DNA <213> Artificial Sequence <223> Description of Artificial Sequence: PCR Primer sequence <400> 120 26 ccagtataag ctgacctttg acaaag <210> 121 <211> 22 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: PCR Primer sequence <400> 121 22 ccaaggtttt agctgtggat ct <210> 122 <211> 24 <212> DNA

<213> Artificial Sequence

<220> <223>	Description of Artificial Sec sequence	quence: PCR	Primer
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<210> <211> <212> <213>	22		
<220> <223>	Description of Artificial Secsequence	quence: PCR	Primer
<400> cacat	123 ttcac actcagctct ga		22
<210> <211> <212> <213>	20		
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<400> cagga	124 gcatt tcgtgaaaga		20
<210> <211> <212> <213>	26		
<220> <223>	Description of Artificial Sec	quence: PCR	Primer
<400> ttttg	125 cacct ttatctgcag cctttg		26
<210> <211> <212> <213>	20		
<220> <223>	Description of Artificial Secsequence	quence: PCR	Primer
<400> tttaa	126 cccga gcttcctcat		20
<210> <211>			

<212> <213>	DNA Artificial Sequence	
<220>	·	
<400> ctgca	· 127 laaatc ttacgacttt gg	22
<210> <211> <212> <213>	· 30	
<220> <223>	Description of Artificial Sequence: PCR Primer sequence	
<400> caaca	· 128 naacaa tggctacatc aaatttagca	30
<210><211><211><212><213>	· 22	
<220> <223>	Description of Artificial Sequence: PCR Primer sequence	
<400> atgac	· 129 cactca gcaaacctga gt	22
<210> <211> <212> <213>	· 22	
<220> <223>	Description of Artificial Sequence: PCR Primer sequence	
<400> ctgca	· 130 waaatc ttacgacttt gg	22
<210> <211> <212> <213>	· 30	
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<400> caaca	- 131 aaacaa tggctacatc aaatttagca	30

	433	COICAZZZ	- ± ; ^\	•	
<210> <211> <212> <213>	22				
<220> <223>	Description of Artificial sequence	Sequence:	PCR	Primer	
<400> atgaca	132 actca gcaaacctga gt			:	22
<210> <211> <212> <213>	22				
<220> <223>	Description of Artificial sequence	Sequence:	PCR	Primer	
<400> ctgcaa	133 aaatc ttacgacttt gg			:	22
<210> <211> <212> <213>	30				
<220> <223>	Description of Artificial sequence	Sequence:	PCR	Primer	
<400> caacaa	134 aacaa tggctacatc aaatttagca	a			30
<210> <211> <212> <213>	21				
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<400> tcagca	135 aaacc tgagtcctgt a				21
<210> <211> <212> <213>	22				
<220> <223>	Description of Artificial sequence	Sequence:	PCR	Primer	
<400> ctgca	136 aaatc ttacgacttt gg	Page	270	:	22

<210> <211> <212> <213>	30		
<220> <223>	Description of Artificial Sequence: sequence	PCR Primer	
<400> caacaa	137 aacaa tggctacatc aaatttagca		30
<210> <211> <212> <213>	22		
<220> <223>	Description of Artificial Sequence: sequence	PCR Primer	
<400> atgaca	138 actca gcaaacctga gt		22
<210> <211> <212> <213>	22		
<220> <223>	Description of Artificial Sequence: sequence	PCR Primer	
<400> ctgca	139 aaatc ttacgacttt <b>gg</b>		22
<210> <211> <212> <213>	30		
<220> <223>	Description of Artificial Sequence: sequence	PCR Primer	
<400> caaca	140 aacaa tggctacatc aaatttagca		30
<210> <211> <212> <213>	22		
<220> <223>	Description of Artificial Sequence: sequence	PCR Primer	

		CONAZZZ	- T - V	F	
<400> atgaca	141 actca gcaaacctga gt				22
<210> <211> <212> <213>	22				
<220> <223>	Description of Artificial sequence	Sequence:	PCR	Primer	
<400> gggcta	142 ataag tcagtcggaa gt				22
<210> <211> <212> <213>	26				
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<400> gagga	145 cagct ttgatttcat tg				22
<210> <211> <212> <213>	27				
<220> <223>	Description of Artificial	Sequence: Page	PCR 272	Primer	

## sequence

<400> tggatt	146 tgat ccatttcctc tctacca	27
<210> <211> <212> <213>	22	
<220> <223>	Description of Artificial Sequence: PCR Primer sequence	
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